SEQUENCE LISTING

Val

<110> MUTABILIS <120> Comprising of polypeptides specific to pathogenic strains and their use as vaccines and in immunotherapy <130> .2209 <160> 160 <170> PatentIn version 3.1 <210> 10 <211> 163 <212> PRT <213> Escherichia coli <400> 1 15 Met Lys Leu Lys Ala Ile Ile Leu Ala Thr Gly Leu Ile Asn Cys Ile 5 10 1 15 20 Val Phe Ser Ala Gln Ala Val Asp Thr Thr Ile Thr Val Thr Gly Aśn 20 25 30 25 Val Leu Gln Arg Thr Cys Asn Val Pro Gly Asn Val Asp Val Ser Leu 35 40 45 30 Gly Asn Leu Tyr Val Ser Asp Phe Pro Asn Ala Gly Ser Gly Ser Pro 50 55 60 35 Trp Val Asn Phe Asp Leu Ser Leu Thr Gly Cys Gln Asn Met Asn Thr 75 65 70 80 40 Val Arg Ala Thr Phe Ser Gly Thr Ala Asp Gly Gln Thr Tyr Tyr Ala 85 90 95 Asn Thr Gly Asn Ala Gly Gly Ile Lys Ile Glu Ile Gln Asp Arg Asp 100 105 110 50 Gly Ser Asn Ala Ser Tyr His Asn Gly Met Phe Lys Thr Leu Asn

115 120 125

Gln Asn Asn Ala Thr Phe Asn Leu Lys Ala Arg Ala Val Ser Lys

130 135 140

Gly Gln Val Thr Pro Gly Asn Ile Ser Ser Val Ile Thr Val Thr

Tyr

145

160

15 Thr Tyr Ala

25 Met Lys Met Thr Arg Leu Tyr Pro Leu Ala Leu Gly Gly Leu Leu . Leu 1 5 10 15

30 Pro Ala Ile Ala Asn Ala Gln Thr Ser Gln Gln Asp Glu Ser Thr
Leu
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25
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35 Val Val Thr Ala Ser Lys Gln Ser Ser Arg Ser Ala Ser Ala Asn Asn 35 40 45

40 Val Ser Ser Thr Val Val Ser Ala Pro Glu Leu Ser Asp Ala Gly Val 50 55 60

45 Thr Ala Ser Asp Lys Leu Pro Arg Val Leu Pro Gly Leu Asn Ile Glu
65 70 75 80

50 Asn Ser Gly Asn Met Leu Phe Ser Thr Ile Ser Leu Arg Gly Val Ser

85 90 . 95

Ser Ala Gln Asp Phe Tyr Asn Pro Ala Val Thr Leu Tyr Val Asp 5 Gly 100 105 110

Val Pro Gln Leu Ser Thr Asn Thr Ile Gln Ala Leu Thr Asp Val

10 Gln

115 120 125

Ser Val Glu Leu Leu Arg Gly Pro Gln Gly Thr Leu Tyr Gly Lys

15 Ser

130 135 140

Ala Gln Gly Gly Ile Ile Asn Ile Val Thr Gln Gln Pro Asp Ser
Thr
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165 170 175

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40 Ala Thr Gly Ser Asp Asp Leu Gly Gly Thr Arg Ala Ser Ile Gly Asn
210 215 220

Val Lys Leu Arg Leu Ala Pro Asp Asp Gln Pro Trp Glu Met Gly Phe
225 230 235
240

•	
	4/370

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	Gln Met	His	Tyr	Ser	Arg	Thr	Phe	Pro	Ser	Gly	Ser	Leu	Ile	Val	Asn
	305 320					310					315	ı			
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	Gly	Asp	Ala	Arg	Thr	Val	Asp	Met	Val	Phe	Gly	Leu	Tyr	Arg	Gln
	Asn			340					345			٠		350	
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	Leu	Ser	Ser	Thr	Gly	Tyr	Thr	Thr	Ala	Glu	Thr	Leu	Ala	Ala	Tyr
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45					-										
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	Δκα	Dha	Sar	Hie	Asp	T.ve	Sar	Sar	Thr	Gln	ጥ _ህ ዮ	Hie	Glv	Ser	Met
	Leu	rne	Der	117.2	_	пуз	DCT	Det	1111		тŸт	1120	OLY	DCT	
5					405					410					4,15
•	Gly	Asn	Pro	Phe	Gly.	Asp	Gln	Gly	Lys	Ser	Asn	Asp	Asp	Gln	Val
	Leu		•	420	_	_	•	_	425			_	_	430	
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	-	Gln	Leu	Ser	Ala	Gly	Tyr	Met	Leu	Thr	Asp	Asp	Trp	Arg	Val
	Tyr	•	435					440					445		
15			•												
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	Thr Asn	Ala	Gly	Leu	Asp	Ala	Lys	Pro	Phe	Val	Ala	GLu	Lys	Ser	Ile
25	465 480					470					475				
4				•	-									•	
	-	Glu	Leu	Gly	Thr	Arg	Tyr	Glu	Thr	Ala	Asp	Val	Thr	Leu	Gĺn
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	Ala Pro	Thr	Phe	Tyr	Thr	His	Thr	Lys	Asp	Met	Gln	Leu	Tyr	Ser	Gly
35				500					505					510	
	TT - 7	a 1	N/	C1	ml	Т	0	71	71 7 ~	C1	T	71 7	7\ a.s.	77 7 0	⊞b zo
	Val Gly	СΤΆ		GIN	THE	Leu	ser		Ald	GTÀ	тйг	Ald		Ala	TIIT
40		i,	515					520					525		
	Val	Glu	Leu	Glu	Ala	Lys	Trp	Arg	Phe	Ala	Pro	Gly	Trp	Ser	Trp
45	Asp	530				_	535					540	-		-
73		330					333		-			310			
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		Val	Asn	Gly	Val	Ile	Asp	Thr	Arg	Tyr	Gly	Ala	Leu	Met	Pro
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	Leu	Ala	Val	Asn	Leu	Val	Gly	Pro	His	Tyr	Phe	Asp	Gly	Asp	Ásn
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	Len	Ara	Gln	Glv	Thr	Tvr	Ala	Thr	Leu	Asp	Ser	Ser	Leu	Glv	Tro
20	Gln	_		JJ		- 11 -	615					620			
	70 7	m)	C 1	70	3.6	70	- 1		77 7	m	** 7	70		-	D1
25.	Ala Asp 625	Thr	GLU	Arg	Met	Asn 630	TTE	ser .	Val	Tyr	Va1 635	Asp	Asn	Leu	Phe
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	Gln	Val	Asn	Met	Gly	Arg	Thr	Val	GLy	Ile	Asn	Thr	Arg	Ile	Asp
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		7/370			
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10	Gly Leu	Lys	Ala 35	Gly	Ala	Ser	Val	Met	Ser	Leu	Ser	Asp	Gln 45	Arg	Phe	
15	Ser His	Gly 50	Asp	Glu	Glu	Glu	Thr 55	Ser	Lys	Tyr	Lys	Glу 6О	Gly	Asp.	Asp.	
20	Asp Pro 65	Thr	Val	Phe	Ser	Gly 70	Gly	Ile	Ala	Val	Gly 75	Tyr	Asp	Phe	Tyr	80
25	Gln Gly	Phe	Ser	Ile	Pro 85	Val	Arg	Thr	Glu	Leu 90	Glu	Phe	Tyr	Ala	Arg 95	
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	Ser Gly	Arg	Ser	Gly	Ser	Ala	Asp	Asn	Phe	Ala	Trp	Ser	Leu	Glу	Ala
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•		Arg	Tyr	Asp	Val	Thr	Pro	Asp	Ile	Ala	Leu	Asp	Leu	Ser	Tyr
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	Tyr Gly	Leu	Asp	Ala	Gly	Asp	Ser	Ser	Val	Ser	Tyr	Lys	Asp	Gl.u	Trp
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	Asp Leu	Lys	Tyr	Lys	Ser	Glu	Val	Asp	Val	Lys	Ser	His	Asp	Ile	Met
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35	Ile 1	1,5			5			200		10	22,	200		,	15
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Thr Val Val Leu Asp Asn Ala Tyr Thr Ser Asp Leu Ile Ala Ala 50 Asn 50 55 60

9/	12

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10	Asn Asn	Val	Asn	Asn	Val	Thr	Ser	Phe	Gly	Gly 90	Thr	Ala	Glu	Asn	Thr 95	
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25	Val Ala	Thr 130	Val	Ser	Asn	Gly	Gln 135	Ala	Thr	Phe	Asn	Leu 140	Lys	Val	Arg	
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	Ala Asp	Ala	Asn	lle	Asp	Ile	Ser	Asn	Val	Trp	Ala	Arg	Asp	Tyr	Leu	
10	-	50					55					60				
	Leu Thr	Ala	Gln	Asn	Lys	Gly	Ile	Phe	Gl'n	Pro	Gly	Ala	Thr	Asp	Val	
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	Ile Ile	Thr	Leu	Lys	Asn	Gly	Asp	Lys	Phe	Ser	Phe	His	Asn	Leu	Ser	
20					85		•	(90	٠				95	
	Pro Gly	Asp	Phe		Gly	Ala	Ala	Ala	Ser	Gly	Ala	Ala	Thr	Ala	Ile	
25				100			٠.		105	,				110		4
	Gly Ala	Ser	Tyr	Ser	Val	Thr	Val	Ala	His	Asn	Lys	Lys	Asn	Pro	Gln	
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	Ala Arg	Glu	Thr	Gln	Val	Tyr	Ala	Gln	Ser	Ser	Tyr	Arg	Val	Val	Asp	-
35		130					135			•		140				
	Arg Val	Asn	Ser	Asn	Asp						Leu		Lys	Phe	Val	
40	145 160	,				150	٠,	•			155		•	•		
45	Glu Ser	Thr	Val	Gly	Ala	Thr	Pro	Ala	Glu	Thr	Asn	Pro	Thr	Thr	Tyr	
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Asp Ala Leu Glu Arg Tyr Gly Ile Val Thr Ser Asp Gly Ser Lys 50 Lys 180 185 190

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	Glu	Ser	Lvs	Tle	Ser	Thr	Asn	Ser	Δla	, Tur	Ser	His	Asn	T.611	T.A.1
	Ser		пуо	110		1111		DCI	ALG	- y -	DGI		дзр	пец	пец
10	•	210					215					220			
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	240								•	•					
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				-											
25	Gly Leu	Ser	Gly	Ala	Tyr	Leu	Tyr	Asp	Asn	Lys	Leu	Glu	Lys	Trp	Val
				260					265					270	
	\7~ 1	C1.	Th x	Thr	nic	Cly	Tlo	7\] =	Sor	77 = 7	Asn	Clv	7 00	Cln	T 011
30	Thr	GTĀ		7117	UTS	GTĀ	T'TE		ser	vaı	ASII	дтλ		GTII	пеп
			275					280			·		285		
		Ile	Thr	Lys	Tyr	Asn	Asp	Lys	Leu	Val	Ser	Glu	Leu	Lys	Asp
35	Thr	290					295					300			
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40	Tyr Asn	Ser	His	Lys	Ile	Asn	Leu	Asn	Gly	Asn	Asn	Val	Thr	Ile	Lys
	305 320					310					315				
	J20 ,								•						
45		Asp	Ile	Thr	Leu	His	Gln	Asn	Asn	Ala	Asp	Thr	Thr	Gly	Thr
	Gln				325					330					335
													٠		
50	Glu Asp	Lys	Ile	Thr	Lys	Asp	Lys	Asp	Ile	Val	Phe	Thr	Asn	Gly	Gly
															-

340 . 345 . 350

Val Leu Phe Lys Asp Asn Leu Asp Phe Gly Ser Gly Gly Ile Ile
5 Phe
355 360 365

Asp Glu Gly His Glu Tyr Asn Ile Asn Gly Gln Gly Phe Thr Phe
10 Lys
370 375 380

Gly Ala Gly Ile Asp Ile Gly Lys Glu Ser Ile Val Asn Trp Asn
15 Ala
385 390 395

20 Leu Tyr Ser Ser Asp Asp Val Leu His Lys Ile Gly Pro Gly Thr Leu 405 410 415

25 Asn Val Gln Lys Lys Gln Gly Ala Asn Ile Lys Ile Gly Glu Gly
Asn
420 425 430

30 Val Ile Leu Asn Glu Glu Gly Thr Phe Asn Asn Ile Tyr Leu Ala Ser 435 440 445

35 Gly Asn Gly Lys Val Ile Leu Asn Lys Asp Asn Ser Leu Gly Asn Asp 450 455 460

40 Gln Tyr Ala Gly Ile Phe Phe Thr Lys Arg Gly Gly Thr Leu Asp Leu 465 470 475

Asn Gly His Asn Gln Thr Phe Thr Arg Ile Ala Ala Thr Asp Asp Gly
485
490
495

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	Ile		515					520					525		
10	T	T	mb		7	· 〒 7 _	7)	Cl	G1	71	T	T	, ml		7 . 7 -
	Lys	ьеи 530	THE	HIS	ASN	тте	535	ser	GTU	Asp	туѕ	ьуs 540	Thr	Asn	Ala
15		330					,					340			
	Leu Ser	Ile	Leu	Asp	Gly	Ser	Val	Asn	Thr	Lys	Asn	Asp	Val	Glu	Val
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2.7	Phe		595		-	٠,		600					605		
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	Asp Asn	Trp	Lys	Thr	Gly	Val.	Phe	Lys	Phe	Asp	Thr	Leu	His	Leu	Asn
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	Ala Asp	Asn	Lys	Ser	Ala	Ile	Thr	Ile	Gly	Asp	Lys	Asn	Val	Тух	Ile
5		•		660					665					67 O	
J	70	_	- T	6 3		<i>.</i>	 1	m1	_	-	6 3	7.		_,	_
	Asn Gln	Leu	Ala	GLY	Lys	Asn	TTe	Thr	Asn	Asn	GTY	Phe	Asp	Phe	Lys
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	Thr	Ile	Ser	Thr	Asn	Leu	Ser	Ile	Glv	Glu	Thr	Lvs	Phe	Th≆	Glv
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15		090					093					700		-	
		Thr	Ala	His	Asn	Ser	Gln	Ile	Ala	Ile	Gly	Asp	Gln	Ala	Val
	Val 705					710					715				
20	720					•									
	Thr	Len	Asn	Glv	Ala	Thr	Phe	Tien	Asn	Asn	Thr	Pro	Tle	Seæ	'T]_
25	Asp	БСС	11011	CIY	725	1111	1110		1100	730	1111	-	110		
23					123					730					735
		Gly	Ala	Lys	Val	Ile	Ala	Gln	Asn	Ser	Met	Phe	Thr	Thr	Lys
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								•	·						
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	_	_													
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TJ	800					150		,			190				
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8.15

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	Glu Thr	Thr	Leu	Leu	Tyr	Gly	Phe	Asp	Thr	Ala	Tyr	Arg	Gly	Ala	Ile
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	Ala Asn	Pro	Lys	Ala	Thr	Val	Ser	Met	Asn	Asn	Ala	Ile	Trp	His	Leu
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-	Q	C3	0	0	T] -	71	77	T	0.1	m.	-	73	~		
	Arg	Gln.	ser	ser	тте		Arg	ьeu	GIU	Thr		Asp _.	ser	Met	Val
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			•		•					•					
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30	Thr Ala	Ile	Asp	Asp	Ser	Ala	Phe	Val	Leu	Arg	Ala	Asn	Leu	Ala	Gln
		-		900				-)	905					910	
,	_	T				_	_	_	_						
35	Asp	Gln	Leu	Val	Val	Asn	Lys	Ser	Leu	Ser	Gly	Lys	Asn	Asn	Leu
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	960														
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965 970 975

															-
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10	Val Gly	Ala	Asn . 995	Ala .	Asp	Ala 2	Ala Li	ys :	Lys :	Ala '	Thr :		eu 1	Met :	Ser
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50	Trp	Tyr 1145	Ala	Gly	Ala	Glu	Val 1150	Gly	Tyr	Arg	Tyr	His 1155	Val	Thr	Asp

5	Ser Ala 1160		e Glu I		ln 1	Ala	Glu	Leu	Val	Tyr 1170	Gly	Ala	Val
	Ser Gly 1175		n Phe S		rp :	Lys	Asp	Gln	Gly	Met 1185	Asn	Leu	Thr
10	Met Lys 1190		s Asp' E		sn : 195	Pro _.	Leu	Ile	Gly	Arg 1200	Thr	Gly	Val
15	Asp Val 1205		s Ser E		er (210	Gly	Lys	Asp	Trp	Lys 1215	Val	Thr	Ala
20	Arg Ala 1220		u Gly T		ln 1 225	Phe ·	Asp	Leu	Phe	Ala 1230	Asn	Gly	Glu
25	Thr Val 1235	Leu Ar	g Asp A		er (240	Gly	Glu	Lys	Arg	Ile 1245	Lys	Gly	Glu
	Lys Asp 1250	Gly Ar	g Met I		et 1 255	Asn	Val	Gly	Leu	Asn 1260	Ala	Glu	Ile
30	Arg Asp 1265	Asn Le	u Arg F		ly 1 270	Leu	Glu	Phe	Glu	Lys 1275	Ser	Ala	Phe
35	Gly Lys 1280	Tyr As	n Val A		sn 2 285	Ala	Ile	Asn	Ala	Asn 1290	Phe	Arg	Tyr
40	Ser Phe 1295				٠								
45	<212> PI	42 RT scheric	nia col	i.									,
50	Met Ile A Ala 1	Asn Ile	Pro Se	er Pro	o Thi	r Al	a Va 10		ıl Me	et Ala	Leu	val 15	-

Ile Ser Thr Leu Pro Ser Pro Ser Arg Val Lys Leu Met Pro Tyr Pro 5 20 25 30 Pro Arg Ala His Asn Thr Thr Gly Leu Leu Pro Val Arg Glu Ile 10 35 40 45 Phe Pro His His Gly Asp Asp Gly Arg Asn Ser Ile Glu Pro Ser 15 50 55 60 Ser Arg Ala Ala His Thr Asp Arg Leu Arg Phe Val Cys Met Thr Arg .80 20 70 75 65 Thr Gly Ser Thr Thr Ser Arg Pro Phe Cys Pro Ile Pro Arg Ser Pro 25 85 90 95 Ala Leu Asn Ala Ser Gly Gln Gln Asp Ser Gly Phe Trp Gly Val Ser 30 100 105 110 Ser Ile Pro Gly Asp Ile Leu Met Phe Gln Leu His Val Leu Ile 120 35 115 125 Phe Ile Cys Lys Ile Asn Leu Ser Asp Asn Asn Ile Ser Tyr · 130 135 140 40 <210> 7 <211> 318 <212> PRT 45 <213> Escherichia coli <400> 7 Met Tyr Ala Arg Glu Tyr Arg Ser Thr Arg Pro His Lys Ala Ile Phe 50 1 5 10 15

	Phe Ala	His	Leu	Ser	Cys	Leu	Thr	Leu	Ile	*Cys	Ser	Ala	Gln	Val	Tyr	
5				20					25					30		
	Lys	Pro	Asp	Met	Arg	Pro	Leu	Gly	Pro	Asn	·Ile	Ala	Asp	Lys	Glv	
	Ser	•	35		_			40	•				45	-	.	
10	,															
	Val Thr		Tyr	His	Phe	Ser		Thr	Ser	Phe	Asp		Val	Asp	Gly	
15		50					55					60				
		His	Tyr	Arg	Val	Trp	Thr	Ala	Val	Pro	Asn	Thr	Thr	Ala	Pro	٠
20	Ala 65					70					75					80
20	Ser	Glv	Tur	Pro	Tle	T.e.ii	Tur	Met	T, <u>011</u>	Asn	Gliv	Asn	Ala	Val	Mat	
	Asp	CTY	- <u>y</u> -	110	. 85	Lou	- y -	1100	пса	90	Gry	71011	ΜΙΔ	vai	95	
25															50	
	Arg Pro	Leu	Asp	Asp	Glu	Leu	Leu	Lys	Gln	Leu	Ser	Glu	Lys	Thr	Pro	
30				100					105					110		
			Val	Ala	Val	Gly	Tyr	Gln	Thr	Asn	Leu	Pro	Phe	Asp	Leu	
2.5	Asn		115				•	120					125			•
35	C	7)	70.7 _	W	7)	F	mla	D	70 T _	7 . 7	0 1.	G .	- T	_	1	
	Asp	-	Ala	Tyr	Asp	Tyr		Pro	Ата	Ата	GLu		Arg	Lys	Thr	,
40		130					135					140				
	Leu Phe	His	Ser	Gly	Arg	Phe	Ser	Arg	Lys	Ser	Gly	Gly	Ser	Asn	Asn	
45	145					150					155					
.5	100											·				
	Arg Leu	Gln	Leu	Leu	Glu	Thr	Arg	Ile	Ala	Pro	Lys	Val	Glu	Gln	Gly	
50					165			٠		170					175	

	Asn Gly	Ile	Asp	Arg	Gln	Arg	Arg	Gly	Leu	Trp	Gly	His	Ser	Tyr	Gly
5	_			180					185					190	
5															
	Leu Tyr	Phe	Val	Leu	Asp	Ser	Trp	Leu	Ser	Ser	Ser	Tyr	Phe	Arg	Ser
10	4		195					200					205		
10															
	Tyr Ser	Ser	Ala	Ser	Pro	Ser	Leu	Gly	Arg	Gly	Tyr	Asp	Ala	Leu	Leu
15	•	210					215		*			220			•
15					· .										
	Arg Ala	Val	Thr	Ala	Val	Glu	Pro	Leu	GIn	Phe	Cys	Thr	Lys	His	Leu
20	225 240					230					235				
20	210								•						
	Ile	Met	Glu	Gly	Ser	Ala	Thr	Gln	Gly	Asp	Asn	Arg	Glu	Thr	His
25	Ala				245				٠	250					255
				•				•		200					200
	Val	Gly	Val	Leu	Ser	Lys	Ile	His	Thr	Thr	Leu	Thr	Ile	Leu	Lys
30	Asp			260					265					270	
		Gly	Val	Asn	Ala	Val	Phe	Trp	Asp	Phe	Pro	Asn	Leu	Gly	His
35	Gly		275					280			•		285		·
		Met	Phe	Asn	Ala	Ser	Phe	Arg	Gln	Ala	Leu	Leu	Asp	Ile	Ser
40	Gly	290					295					300			
			,												
		Asn	Ala	Asn	Tyr		Ala	Gly	Cys	His		Leu	Ser	His	
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	<211	.> 7	25												
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	21/3

	Mot	A ra	Tle	Asn	Lvs	Ile	Leu	Trp	Ser	Leu	Thr	Va l	Leu	Leu	Val	
	Gly	7119	110	11011	5			-		10					15	
5	1 .				J							_				
	Leu	Asn	Ser	Gln	Val	Ser	Val	Ala	Lys	Tyr	Ser	Asp	Asp	Asp	Asn	
	Asp			20					25					30 ·		
10										_				-	a 1	
	Glu Gln	Thr	Leu	Val	Val	Glu	Ala	Thr	Ala	Glu	Gln	Val		ьуs	GIN	
15			35					40					45			
13	Drác	C] 11	77 a 1	Ser	Val	Tle	Thr	Ser	Glu	Asp	Ile	Lys	Lys	Thr	Pro	
	Pro			001			55			_		60				
20		50	•													
	Val	Asn	Asp	Leu	Ser	Asp	Ile	Ile	Arg	Lys	Met	Pro	Gly	Val	Asn	
	Leu 65					70			•		75					80
25											-	77	7)	Cl ~	Tlo	
	Thr Asp		Asr	ser	Ala	Ser	Gly	Thr	Arg		Asn	Asn	Arg	GT11		
30					85					90					95	
50	Tle	a Arc	r Gly	, Met	: Gly	Pro	Glu	ı Asn	Thr	Leu	ı Ile	. Leu	Ile	Asp	Gly	
	Val		,	100					105					110		
35				100	,											
			l Th	r Sei	r Arg	Asn	se:	r Val	Arc	ј Туг	Ser	r Txp	Arg	g Gly	Glu	
	Ar	g	11	5				120)				125	5		
40						1			- 17-	l Dwa	n Dro	പ്പ	ı Gla	, val	Glu	
	As; Ar		r Ar	g Gl	y Asp	Thi			o va.	r bro) LT	Gli		ı val	. 010	•
45		13	0			•	13	5				1.40	J			
		e Gl	u Va	1 Il	e Aro	g Gl	y Pr	o Al	a Ala	a Ala	a Ar	g Ty:	r Gl	y Se:	r Gly	7
	Al 14	a				15					15					
50																

															,
	Ala Trp	Gly	Gly	Val	Val	Asn	Ile	Ile	Thr	Lys	Arg	Pro	Thr	Asn	Asp
5	TTP				165	-				170					175
	His Glu	Gly	Ser	Leu	Ser	Leu	Tyr	Thr	Asn	Gln	Pro	Glu	Ser	Ser	Glu
10	OLU			180					185					190	
	_	Ala.	Thr	Arg	Arg	Ala	Asn	Phe	Ser	Leu	Ser	Gly	Pro	Leu	Ala ^{'.}
15	Gly	•	195					200				,	205		
	Asp Ala	Ala	Leu	Thr	Thr	Arg	Leu	Tyr	Gly	Asn	Leu	Asn	Lys	Thr	Asp
20	ALA	210					215					220	,		
	Asp Gly	Ser	Trp	Asp	Ile	Asn	Ser	Pro	Val	Gly	Thr	Lys	Asn	Ala	Ala
25	225					230′					235				
	His	Glu	Gly	Val	Arq	Asn	Lys	Asp	Ile	Asn	Gly	Val	Val	Ser	Trp
30	Lys		_		245		-	*		250	_				255
	Leu Gln	Asn	Pro		Gln	Ile	Leu	Asp		Glu	Val	Gly	Tyr		Arg
35				260		-			265					.270	
	_	Asn	Ile	Tyr	Ala	Gly	Asp	Thr	Gln	Asn	Ser	Ser	Ser	Ser	Ala
40	Val		275					280					285		
	Thr	Glu	Ser	T.e.11	Δla	T.vs	Ser	Glv	Lvs	Glų	Thr	Asn	Ara	T.e11	Tur
45	Arg	290	DCI	ПСИ	71110	11 1 1 1	295		Lyo	0 ± 4		300	**** 9	Lou	- <i>y</i> -
15		2,70			•							200			
50	Gln Ser 305 320	Asn	Tyr	Gly	Ile	Thr 310	His	Asn	Gly	Ile	Trp 315	Asp	Trp	Gly	Gln

5	Arg Glu	Phe	Gly	Val	Tyr 325	Tyr	Glu	Lys	Thr	Asn 330	Asn	Thr	Arg	Met	Asn 335
10	Gly Phe	Leu	Ser	Gly 340	Gly	Gly	Gļu	Gly	Arg	Ile	Leu	Ala	Gly	Glu 350	Lys
15	Thr Ile	Thr	Asn 355		Leu	Ser	Ser	Trp 360	Arg	Thr	Ser	Gly	Glu 365	Leu	Asn
20	Pro Trp	Leu 370	Asn	Val	Met	Val	Asp 375	Gln	Thr	Leu _.	Thr	Val 380	Gly	Ala	Glu
25	Asn Asn 385 400	Arg	Asp	Lys	Leu	Asp	Asp	Pro	Ser	Ser	Thr 395	Ser	Leu	Thr	Val
30	Asp Ser	Arg	Asp	Ile	Ser	Gly	Ile	Ser	Gly	Ser 410	Ala	Ala	Asp	Arg	Ser
35	Lys Glu	Asn	His	Ser 420	Gln	Ile	Ser	Ala	Leu 425	Tyr	Ile	Glu	Asp	Asn 430	Ile
40	Pro Leu	Val	Pro 435	Gly	Thr	Asn	Ile	Ile 440		Glý	Leu	Arg	Phe	Asp	Tyr
45	Ser Glu	Asp 450	Ser	Gly	Gly		Phe	Ser	Pro	Ser	Leu	Asn 460	Leu	Ser	Gln
50	Leu Lys	Gly	Asp	Tyr	Phe	Lys	Val	Lys	Ala	Gly	Val	Ala	Arg	Thr	Phe

465 470 475

480

5 Ala Pro Asn Leu Tyr Gln Ser Ser Glu Gly Tyr Leu Leu Tyr Ser Lys

485 490 495

10 Gly Asn Gly Cys Pro Lys Asp Ile Thr Ser Gly Gly Cys Tyr Leu Ile 500 505 510

15 Gly Asn Lys Asp Leu Asp Pro Glu Ile Ser Val Asn Lys Glu Ile Gly 515 520 525

20 Leu Glu Phe Thr Trp Glu Asp Tyr His Ala Ser Val Thr Tyr Phe Arg 530 535 540

25 Asn Asp Tyr Gln Asn Lys Ile Val Ala Gly Asp Asn Val Ile Gly Gln 545 550 550

30
Thr Ala Ser Gly Ala Tyr Ile Leu Lys Trp Gln Asn Gly Gly Lys Ala
565
570
575

Leu Val Asp Gly Ile Glu Ala Ser Met Ser Phe Pro Leu Val Lys Glu
580 585 590

40
Arg Leu Asn Trp Asn Thr Asn Ala Thr Trp Met Ile Thr Ser Glu Gln
595
600
605

Lys Asp Thr Gly Asn Pro Leu Ser Val Ile Pro Lys Tyr Thr Ile
Asn
610
615
620

	W	O 2005	5/09782	3									PCT/E	P2005	/003972
								25/37							
	Asn Asn 625	Ser	Leu	Asn	Trp	Thr 630	Ile	Thr	Gln	Ala	Phe 635	Ser	Ala	Ser	Phe ,
5	640					050					033				
	Trp Arg	Thr	Leu	Tyr	Gly	Arg	Gln	Lys	Pro	Arg	Thr	His	Ala	Glu	Thr
10	1119				645					650					655
	Ser Ser	Glu	Asp	Thr	Gly	Gly	Leu	Ser	Gly	Lys	Glu	Leu	Gly	Ala	Tyr
15				660					665				•	670	
	Leu Leu	Val	Gly	Thr	Asn	Phe	Asn	Tyr	Asp	Ile	Asn	Lys	Asn	Leu	Arg
20			675					680					685		
	Asn Ser		Gly	Val	Ser	Asn		Leu	Asn	Lys	Gln	•	Phe	Arg	Ser
25		690					695					700			
	Gly	Gly	Ala	Asn	Thr	Tyr	Asn	Glu	Pro	Gly		Ala	Tyr	Tyr	Ala [.]
30	705					710					715	•			u.
	Val	Thr	Ala	Ser	Phe 725										
35															
40	<210 <211 <212 <213 <400	L>) L014 PRT Esche	erich	nia d	coli		•						,	
				Gln	Trp	Gln	Gln	Lys	Tyr	Leu	Leu	Glu	Tyr	Asn	Glu
45	Leu 1				5					10					15
	Val	Ser	Asn	Phe	Pro	Ser	Pro	Glu	Arg	Val	Val	Ser	Asp	Tyr	Ile

Lys

	Asn Asp	Cys	Phe	Lys	Thr	Asp	Leu	Pro	Trp	Phe	Ser	Arg	Ile	Asp	Pro	
5	2101		35					40					45			
J	Asn	Ala	Tvr	Phe	Ile	Cvs	Phe	Ser	Gln	Asn	Ara	Ser	Asn	Ser	Ara	
,	Ser	50	-1-			-1-	55				9	60	11011		9	
10											•					
	Tyr Thr	Thr	Gly	Trp	Asp	His	Leu	Gly	Lys	Tyr	Lys	Thr	Glu	Val	Leu	
15	65					70				•	75					80
13	Tou	Th w	Cln	7.] -	7.] -	Tou	Tlo	7 9 7	Tlo	Cler	П	71 22 22	Dha	71 ~	777	-
	Phe.		GIII	Ата		ьеи	тте	ASII	TTE	Gly	т Ат	Arg	rne	ASP		
20	ē				85				•	90				•	95	
-		Asp	Ala	Asn	Ser	Ser	Thr	Gly	Ile	Tyr	Lys	Thr	Lys	Ser	Ala	
2.5	Asp			100					105		•			110		
25																
	Val Leu	Phe		Glu	Glu	Asn	Glu		Lys	Met	Leu	Pro		Glu	Tyr	
30			115					120					125			
	His	Phe	Leu	Gln	Lys	Cys	Asp	Phe	Ala	Gly	Val	Tyr	Gly	Lys	Thr	
	Leu	130					135				· v	140				
35			•													,
	Ser Lys	Asp	Tyr	Trp	Ser	Lys	Tyr	Tyr	Asp	ras.	Phe	Lys	Leų	Leu	Leu	
40	145 160					150				•	155					•
														•		
	Asn Leu	Tyr	Tyr	Ile	Ser	Ser	Ala	Ĺeu	Tyr	Leu	Tyr	Lys	Asn	Gly	Glu	
45	100				165					170					175	
	7\ c.r.	Glu	Δνα	Glu	Туу	Aen	Dhe	Ser	Me+	Aen	70.7 =	T.e.i	Δερ	Δκα	962	
50	Asp	. Gru	чтд		т Л т	VOII	E II G	DET		Asn	ALA	пeп	UDII	_	೧೯۲	
50				180					185					190		

	Asn Asp	Ile	Ser	Leu	Leu	Phe	Phe	Asp	Ile	Tyr	Gly	Tyr	Tyr	Ala	Ser
5			195					200					205		
	Ile Gly		Val	Ala	Lys	Asn		Asp	Lys	Val	Met		Phe	Ile	Pro
10		210					215					220			:
	Leu	Lys	Lys	Pro	Phe		Phe	Lys	Lys	Asn		Ala	Asp	Leu	Arg
15	225 240					230	-				235				
		Leu	Lys	Glu	Leu	Ile	Lys	Asp	Ser	Asp	Asn	Lys	Gln	Leu	Leu
20	Ser				245					250					255
	Gln Gly	His	Phe	Ser	Leu	Tyr	Ser	Arg	Gln	Asp	Gly	Val	Ser	Tyr	Ala
25	1			260					265			•		270	
	Val. Glu	Asn	Ser	Val	Leu	His	Ala	Ile	Glu	Asn	Asp	Gly	Asn	Phe	Asn
30			275					280					285		•
	Ser Phe	Tyr	Phe	Leu	Tyr	Ser	Asn	Lys	Thr	Leu	Ser	Asn	Lys	Asp	Val
35		290					295					300			•
	Asp Asp	Ala	Ile	Ala	Ile	Ser	Val	Lys	Lys	Arg	Ser	Phe	Ser	Asp	Gly
40	305 320					310					315				
		Val	Ile	Lys	Ser	Asn	Ser	Glu	Ala	Gln	Arg	Asp	Tyr	Ala	Leu
45	Thr				3,25					330					335
50		Leu	Glņ	Thr	Ile	Leu	Ser	Met	Thr	Pro	Ile	Phe	Asp	İle	Val
50	Val			240					215					250	

340

						-									
	Pro Met	Glu	Val	Ser	Val	Pro	Leu	Gly	Leu	Gly	Ile	Ile	Thr	Ser	Ser
5	1100		355					360					365		
	Glv	Ile	Ser	Phe	asp	Gln	Leu	Ile	Asn	Gly	Asp	Thr	Tyr	Glu	Glu
10	Arg	370				•	375			2 - 2		380			
10		0,0				•									
	Arg Leu	Ser	Ala	Ile	Pro	Gly	Leu	Ala	Thr	Asn	Ala	Val	Leu	Leu	Gly
15	385 400					390				-	395				
	400		,												
20	Ser Glu	Phe	Ala	Ile	Pro	Leu	Leu	Ile	Ser	Lys	Ala	Gly	Ile	Asn	Gln.
20	Gra	٠			405	•				410					415
	V > 1	T. 611	Sar	Sar	[= 17	Tla	Aan	Zen	Glu	Gly	Δrα	πhг	T. (21)	Aen	Glu
25	Thr	пеп	per	420	val	116	ASII	ASII	425	Gry	ALG	1111	пец	430	Giu
				420					423						
20		Ile	Asp	Ile	Phe	Leu	Lys	Glu	Tyr	Gly	Ile	Ala	Glu	Asp	Ser
30	Ile		435					440					445		
	0	C	шъ.	7)	T 0	T 0	7		T	·T 0.11	T		, C	C1	C1
35	His		TUŁ	ASII	ьеи	ьеи	_	Val	туз	·Leu·	гуз		ser	дту	GIII
		450					455					460			
40		Asn	Ile	Val	Lys	Leu	Ser	Asp	Glu	Asp	Asn	Gln	Ile	Val	Ala
40	Val 465			144		470					475				
	480														
45	_	Gly	Ser	Ser	Leu	Ser	Gly	I,le	Tyr	Tyr	Glu	Val	Asp	Ile	Glu
	Thr				4.85					490					495
. .		_				_	_	_					~ -		_
50	Gly Asn	Tyr	Glu	Ile	Leu	Ser	Arg	Arg	Ile	Tyr	Arg	Thr	Glu	Tyr	Asn

500 505 510.

Glu Ile Leu Trp Thr Arg Gly Gly Gly Leu Lys Gly Gly Gln Pro
5 Phe 515 520 525

Asp Phe Glu Ser Leu Asn Ile Pro Val Phe Phe Lys Asp Glu Pro 10 Tyr 530 535 540

Ser Ala Val Thr Gly Ser Pro Leu Ser Phe Ile Asn Asp Asp Ser

Ser
545
560

20 Leu Leu Tyr Pro Asp Thr Asn Pro Lys Leu Pro Gln Pro Thr Ser Glu 565 570 575

25 Met Asp Ile Val Asn Tyr Val Lys Gly Ser Gly Ser Phe Gly Asp Arg 580 585 590

30 Phe Val Thr Leu Met Arg Gly Ala Thr Glu Glu Glu Ala Trp Asn Ile
595 600 605

35 Ala Ser Tyr His Thr Ala Gly Gly Ser Thr Glu Glu Leu His Glu Ile
610 615 620

40 Leu Leu Gly Gln Gly Pro Gln Ser Ser Leu Gly Phe Thr Glu Tyr
Thr
625 630 635

Ser Asn Val Asn Ser Ala Asp Ala Ala Ser Arg Arg His Phe Leu Val
645 650 655

30/370

	Val Tyr	Ile	Lys	Val	His	Val	Lys	Tyr	Ile	Thr	Asn	Asn	Asn	Val	Ser
	7			660					665					670	
5		Asn	His	Trp	Ala	Ile	Pro	Asp	Glu	Ala	Pro	Val	Glu	Val	Leu
	Ala		675					680					685		
10	Val	Val	Asp	Ara	Ara	Phe	Asn	Phe	Pro	Glu	Pro	Ser	Thr	Pro	Pro
	Asp	690	-	J			695					700			
15	Ile	Ser	Thr	Ile	Ara	Lvs	Leu	Leu	Ser	Leu	Ara	Tvr	Phe	Lys	Glu
	Ser 705 720				,	710				,	715	-		. <u>.</u> .	
20													•		
	Ile Asn	Glu	Ser	Thr	Ser	Lys	Ser	Asn	Phe	Gln	Lys	Leu	Ser	Arg	Gly
25					725					730					735
	Ile Arg	Asp	Val	Leu	Lys	Gly	Arg	Gly	Ser	Ile	Ser	Ser	Thr	Arg	Gln
30	9			740					745	÷				750	2
	Ala Pro	Ile	Tyr	Pro	Tyr	Phe	Glu	Ala	Ala	Asn	Ala	Asp.	Glu	Gln	Gln
35			755					760					765		
		Phe	Phe	Tyr	Ile	Lys	Lys	Asp	Arg	Phe	Asp	Asn	His	Gly	Ţyr
40	Asp	770				·	775					780			
	Gln Leu	Tyr	Phe	Tyr	Asp	Asn	Thr	Val	Gly	Leu	Asn	Gly	Ile	Pro	Thr
45	785 800					790					795				
	Asn	Thr	Tyr	Thr	Gly	Glu	Ile	Pro	Ser	Asp	Ser	Ser	Ser	Leu	Gly
50	Ser				805		-			810					815

VV C 2003/07/023		1 C 1/E1 2005/0057/2
	31/370	

	Thr Arg	Tyr	Trp	Lys	Lys	Tyr	Asn	Leu	Thr	Asn ·	Glu	Thr	Ser	Ile	Ile
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	Val Glu	Ser	Asn 835	Ser	Ala	Arg	Gly	Ala 840	Asn	Gly	Ile	Lys	Ile 845	Ala	Leu
10	Glu Ser	Val	Gln	Glu	Gly	Lys	Pro	Val	Ile	lle	Thr	Ser	Gly	Asn	Leu
15		850					855					860			
20	Gly Val 865 880	Cys	Thr	Thr	Ile	Val 870	Ala	Arg	Lys	Glu	Gly 875	Tyr	Ile	Tyr	Lys
25	His Gly	Thr	Gly	Thr	Thr 885	Lys	Ser	Leu	Ala	Gly 890	Phe	Thr	Ser	Thr	Thr 895
30	Val Ile	Lys	Lys	Ala 900	Val	Glu	Val	Leu	Glu 905	Leu	Leu	Thr	Lys	Glu 910	Pro
35	Pro Leu	Arg	Val 915	Glu	Gly	Ile	Met	Ser 920	Asn	Asp	Phe	Leu	Val 925	Asp	Tyr
40	Ser Lys	Glu 930	Asn	Phe	Glu	Asp	Ser 935	Leu	Ile	Thr	Tyr	Ser 940	Ser	Ser	Glu
45	Lys Phe 945 960	Pro	Asp	Ser	Gln	Ile 950	Thr	Ile	Ile	Arg	Asp 955	Asn	Val	Ser	Val
50	Pro Ala	Tyr	Phe	Leu	Asp 965	Asn	Ile.	Pro	Glu	His	Gly	Phe	Gly	Thr	Ser 975

	Thr	Val	Leu	Val	Ara	Val.	Asp	Glv	Asn	Val	Val	Val	Ara	Ser	Leu	
5	Ser		2.00	980	1129		110F	<i></i>	985	• • • • • • • • • • • • • • • • • • • •	. • • • •		11119	990	10u	
10	Glu Lys	Ser	_	Ser	Leu	Asn	Ala	-		a Sei	Glı	ı Ile			al Le	eu
10		•	995				•	1000	J				100)5		
15	Val	Phe 1010		. Lys	s Lys	s Phe	e			•						
20	<210 <211 <212 <213 <400	> 4 > E > E	.0 54 RT sche	erich	nia (coli			•		•			-	,	
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	Gln Ala	Thr	Asp	Val	Leu	Val	Ile	Gly		Gly	Pro	Ala	Gly		Ser	
.30				20	. *,				25					30		
a	Ala Arg	Ile		Ala	Gly	Arg	Leu	_	Ala	Arg	Thr	Met		Val	Glu	
35			35					40					45		·	
40	Phe	Gly 50	Ser _.	Leu	Gly	Gly	Val	Leu	Thr	Gln	Val	Gly 60	Val	Glu	Ser	
45	Ala Arg 65	Trp	Tyr	Arg	His	Pro	Gly ·	Thr	Glu	Asp	Cys 75	Glu	Gly	Ile	Cys	80

Glu Tyr Glu Gly Arg Ala Arg Ala Leu Gly Phe Thr Arg Pro Glu Pro 850 85 90 95

														•	
	Gln Ala	Ser	Ile	Ser	Glu	Val	Ile	Asp	Thr	Glu	Gly	Phe	Lys	Val	Val
5	nia			100					105					110	
	Asp	Gln	Met	Ile	Thr	Glu	Ser	Gly	Val	Glu	Pro	Leu	Tyr	His	Ser
	Trp		115					120					125		
10															
	Val Val	Val	Asp	Val	Ile	Lys	Asp	Gly	Asp	Thr	Leu	Cys	Gly	Val	Ile
15	Val	130					135					140			
		Àsn	Lys	Ser	Gly	Arg	Gly	Ala	Ile [.]	Leu	Ala	Lys	Arg	Ile	Val
20	Asp 145					150					155				м
20	160					2			,						
		Thr	Gly	Asp	Ala	Asp	Ile	Ala	Ala	Arg	Ala	Gly	Ala	Pro	Trp
25	Thr			-	165					170					175
											•				
	Lys _. Cys	Arg	Ser	Lys	Asp	Gln	Leu	Met	Gly	Val	Thr	Val	Met	Phe	Ser
30				180					185					190	
		Gly	Val	Asp	Val	Ala	Arg	Phe	Asn	Arg	Phe	Val	Ala	Glu	Glu
35	Leu		195					200				*	205.		
	Lys Thr	Pro	Thr	Tyr	Ala	Asp	Trp	Gly	Lys	Asn	Trp	Thr	Ile	Gln	Thr
40		210					215					220			
	Gly	Lys	Glu	Asp	Pro	Met	Phe	Ser	Pro	Tyr	Met	Glu	Asp	Ile	Phe
45	Thr 225					230					235				
	240	-											-		
	Ara	Ala	Gln	Gln	qsA	Gly	Val	Ile	Pro	Glv	asA	Ala	Gln	Ala	Ile
50	Ala				245					250	**				055

255

		Thr	Trp	Ser	Thr	Phe	Ser	Glu	Ser	Gly	Glu	Ala	Phe	Gln	Met
5	Asn			260					265					270	
	Met	Val	Tyr	Ala	Phe	Gly	Phe	Asp	Cys	Thr	Asp	Val	Phe	Asp	Leu
	Thr					_									
10			275					280					285		
	_	Ala	Glu	Ile	Ala	Gly	Arg	Gln	Gln	Ala	Leu	Trp	Ala	Ile	Asp
15	Ala	290					295					300			
		•										•		ı	
,		Arg	His	Tyr	Val	Pro	Gly	Phe	Glu	Asn	Val	Arg	Leu	Arg	Asn
20.	Phe 305					310			•		315				-
	320							•							
	Glv	Ala	Thr	Leu	Gly	Thr	Arg	Glu	Ser	Arg	Leu	Ile	Glu	Gly	Glu
25	Ile									330					335
					325					330					333
	Arg	Ile	Ala	Asp	Asp	Tyr	Val	Leu	Asn	Gln	Gly	Arg	Cys	Ser	Asp
30	Ser			340					345					350	
2.5		Gly	Ile	Phe	Pro	Glu	Phe	Ile	Asp	Gly	Ser	Gly	Tyr	Leu	·Ile
35	Leu		355					360					365		•
40	Pro Pro	Thr	Thr	Gly	Arg	Phe	Phe	Gln	Ile	Pro	Туг	Gly	Cys	Leu	Val
-10	110	370					375	•				380			•
										~ 7	-		.	G	70.7
45	Gln Gly		Val	Glu	Asn	Leu	Leu	Val	Ala	GTĀ	Arg	Cys	тте	ser	Ala
	385 400					390					395				
50	Val	Val	Ala	His	Thr	Ser	Met	Arg	Asn	. Met	Met	. Cys	Cys	Ala	Val

Thr

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Gly Glu Ala Ala Gly Thr Ala Ala Val Val Ser Leu Gln Gln Asn Cys Thr Val Arg Gln Val Ala Ile Pro Asp Leu Gln Asn Thr Leu Gln Gln Gln Gly Val Arg Leu Ala <210> <211> <212> PRT Escherichia coli <213> <400> Met Ser Ala Lys Arg Arg Leu Leu Ile Ala Cys Thr Leu Ile Thr Ala Ile Tyr His Phe Pro Ala Tyr Ser Ser Leu Glu Tyr Lys Gly Thr Phe Gly Ser Ile Asn Ala Gly Tyr Ala Asp Trp Asn Ser Gly Phe Val Asn Thr His Arg Gly Glu Val Trp Lys Val Thr Ala Asp Phe Gly Val Asn Phe Lys Glu Ala Glu Phe Tyr Ser Phe Tyr Glu Ser Asn Val Leu Asn

His Ala Val Ala Gly Arg Asn His Thr Val Ser Ala Met Thr His

Val

	Arg Gln	Leu	Phe	Asp	Ser	Asp	Met	Thr	Phe	Phe	Gly	Lys	Ile	Tyr	Gly
5				100					105			•		110	
	_	Asp	Asn	Ser	Trp	Gly	Asp	Asp	Leu	Asp	Met	Phe	Tyr	Gly	Phe
10	Gly		115				•	120					125	· <u>.</u>	
	_	Leu	Gly	Trp	Asn	Gly	Glu	Trp	Gly	Phe	Phe	Lys	Pro	Tyr	Ile
15	Gly	130					135					140			
	Leu	His	Asn	Gln	Ser	Glv	Asp	Tvr	Val	Ser	Ala	Lys	Tvr	Gly	Gln
20	Thr 145					150	<u>F</u>	- 2			155		J	1	
	160								,						
25	Asn Phe	Gly	Trp	Asn	Gly	Tyr	Val	Val	Gly	Trp	Thr	Ala	Val	Leu	Pro
23	rne				165					170				•	175
	Thr	Leu	Phe	Asp	Glu	Lys	Phe	Val	Leu	Ser	Asn	Trp	Asn	Glu	Ile
30	Glu			180					185					190	
	Tou	7) en	7\ ~ ~	7) en	7\en	7.1 =	ጥህድ	Thr	Glu	Gln	Gln	Phe	Glv	Arg	Asn
35	Gly	Asp	195	ASII	ASP	ALA	ıyı	200	GIU	CLII	OTH	1110	205	-	71511
								,				,			
40	Leu Lys	Asn	Gly	Gly	Leu	Thr		Ala	Trp	Lys	Phe		Pro	Arg	Trp
		210					215		٠			220	-8		
45		Ser	Val	Thr	Trp	Arg	Tyr	Phe	Asp	Asn	Lys	Leu	Gly	Tyr	Asp
43	Gly 225 240					230					235				
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15	Pro Met	Ala	Leu	Ala 20	Ala	Glu	Ala	Lys	Gln 25	Pro	Asn	Leu	Val	Ile	Ile			
20	Ala Ile	Asp	Asp 35	Leu	Gly	Tyr	['] Gly	Asp	Leu	Ala	Thr	Tyr	Gly 45	His	Gln			
25	Val Phe	Lys	Thr	Pro	Asn	Ile	Asp 55	Arg	Leu	Ala	Gln	Glu 60	Gly	Val	Ļys			
30	Thr Gly 65	Asp	Tyr	Tyr		Pro 70	Ala	Pro	Leu	Ser	Ser 75	Pro	Ser	Arg	Ala	80		
35	Leu Ile	Leu	Thr	Gly	Arg 85	Met	Pro	Phe	Arg	Thr 90	Gly	Ile	Arg	Ser	Trp			
40	Pro Ala		Gly	Lys 100	Asp	Val	Ala	Leu	Gly 105	Arg	Asn	Glu	Leu	Thr	Ile			
45	Leu	Leu	Leu 115	Lys		Gln	Gly	Tyr 120	Asp	Thr	Ala	Met	Met 125	Gly	Lys			
	His	Leu	Asn	Ala	Gly	Gly	Asp	Arg	Thr	Asp	Gln	Pro	Gln	Ala	Gln			

140

50

Asp

							•								
5	Met Asp 145	Gly	Phe	Asp	Tyr	Ser 150	Leu	Ala	Asn	Thr	Ala 155	Gly	Phe	Val	Thr
3	160		.0			100					100			_	
10	Ala Tyr	Thr	Leu	Asp	Asn	Ala	Lys	Glu	Arg	Pro	Arg	Tyr	Gly	Met	Val
		,			165					170					175
15	Pro Lys	Thr	Gly	Trp	Leu	Arg	Asn	Gly	Gln	Pro	Thr	Pro	Arg	Ala	Asp
	-			180					185					190	
20	Met Asn	Ser	Gly	Glu	Tyr	Val	Ser	Ser	Glu	Val	∀al	Asn	Trp	Leu	Asp
20	71311		195					200					205		
25	Lys Val	Lys	Asp	Ser	Lys	Pro	Phe	Phe	Leu	Tyr	Val	Ala	Phe	Thr.	Glu
		210					215					220		,	
30	His Gln	Ser	Pro	Leu	Ala	Ser	Pro	Lys	Lys	Tyr	Leu	Asp	Met	Tyr	Ser
30	225 240			,		230		•			235				
35	_	Met	Ser	Ala	Tyr	Gln	Lys	Gln	His	Pro	Asp	Leu	Phe	Tyr	Gly
	Asp				245					250					255
40	_	Ala	Asp	Lys	Pro	Trp	Arg	Gly	Val	Gly	Glu	Tyr	Tyr	Ala	Asn
	Ile			260					265					270	
45		Tyr	Leu	Asp	Ala	Gln	Val	Gly	Lys	Val	Leu	Asp	Lys	Ile	Lys
	Ala		275					280					285		
50	Met Gly	Gly	Glu	Glu	Asp	Asn	Thr	Ile	Val	Ile	Phe	Thr	Ser	Asp	Asn

290	295	300

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10	Glu Gly	Thr	Asp	Gly	Leu 325	Arg	Gly	Arg	Lys	Asp 330	Asn	Leu	Trp	Glu	Gly 335
15	Ile Gly	Arg	Val	Pro 340	Ala	Ile	Ile	Lys ·	Tyr 345	Gly	Lys	His	Leu	Pro 350	Gln
20	Met Leu	Val	Ser	Asp	Thr	Pro	Val	Tyr 360	Gly	Leu	Asp	Trp	Met 365	Pro	Thr
25	Ala Gly	Lys 370	Met	Met	Asn	Phe	Lys 375	Leu	Pro	Thr	Asp	Arg 380	Thr	Phe	Asp
30	Glu Lys 385 400	Ser	Leu	Val	Pro	Val 390	Leu	Glu	Gln	Lys	Ala 395	Leu	Lys	Arg	Glu
	Pro Asp	Leu	Ile	Phe	Gly 405	Ile	Asp	Met	Pro	Phe 410	Gln	Asp	Asp	Pro	Thr 415
40	Glu Asn	Tŕp	Ala	Ile 420	Arg	Asp	Gly	Asp	Trp 425	Lys	Met	Ile	Ile	Asp 430	Arg
45	Asn Thr	Lys	Pro 435	Lys	Tyr	Leu	Tyr	Asn	Leu	Lys	Ser	Asp	Arg	Tyr	Glu

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	W	2 005	/09782	3				40/37	0				PCT/E	P2005/	003972	2
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	OTY	450					455					460				
5	-	Phe	Leu	Lys	Tyr	Lys	Thr	Asp	Ile	Asp	Asn	Asp	Ser	Leu	Met	
	Lys 465 480					470					475					
10										,		,				
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15	Z010		1 2												ý.	
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20	<213 <400		iscne L3	erici	nia c	COTI										
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	Ile	Ala	Lys	Gly	Phe	Arg	Asn	Glu	Arg	Gly	Phe	Val	Thr	Thr	Thr	
	Ile			20	•				25					30		
30																
	Cys His	Ala	Met	Gly	Glu	Leu	Leu	Ala	Glu	Phe	Leu	Ser	Arg	Asn	Pro	
35			35					40					45			
	Gln	Lvs	Phe	Thr		Pro	Gly	Glu	Phe	Ile	Gly	Pro	Phe	Pro	Ser	
	Gly	50					55				_	60				
40											•					
	Ala Ala	Pro	Ala	Ile	Phe	Ala	Ala	Gln	Val	Ala	Lys	Leu	Ser	His	Arg	
45	65		-			70					75					80
		Phe	Phe	Gly	Cys	Val	Gly	Asn	Asp	Asp	Phe	Ala	Arg	Leu	Ile	
	Ile				85					90					95	

W	O 2005	5/09782	3				41/3	70				PCT/E	EP2005	/003972
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Met			100					105					110	
	Asn	·Ala	Val	Thr	Gly	Thr	Ala	Phe	Val	Ser	Tyr	Gln	Asn	Pro
Gln		115					120		-			125		
Gln Phe	Arg	Asp	Phe	Val	Phe	Asn	Ile	Pro	Asn	Ser	Ala	Cys	Gly	Leu
rne	130					135					140			
Thr Leu	Ala	Glu	His	Ile	Asp	Lys	Asp	Leu	Leu	Lys	Gln	Cys	Asn	His
145 160					1.20				9.	155		,		
	Ile	Val	Gly	Ser	Ser	Leu	Phe	Ser	Phe	Arg	Met	Iļle	Asp	Val
Met				165					1.70	,				175
_	Lys	Ala	Ile	Thr	Thr	Ile	Lys	Ser	Ala	Gly	Gly	Thr	Val	Ser
Phe			180	•				185					190	
Asp Gln	Pro	Asn	Ile	Arg	Lys	Glu	Met	Leu	Ser	Ile	Pro	Glu	Met	Ala
U.J.11		195					200					205	٠	
Ala Glu	Leu	Asp	Tyr	Leu	Ile	Glu	Tyr	Thr	Asp	Ile	Phe	Ile	Pro	Ser
	210			•		215					220		•	
Ser	Glu	Leu	Pro	Phe	Phe	Ala	Arq	His	Lys	Asn	Leu	Ser	Glu	Glu

Ser Glu Leu Pro Phe Phe Ala Arg His Lys Asn Leu Ser Glu Glu Gln 225 230 235

Ile Val Ser Asp Leu Leu His Gly Gly Val Lys His Val Ala Ile Lys 50 245 245 250 255

42/370

Arg Ala Gln Arg Gly Ala Ser Tyr Tyr Lys Leu Lys Asn Gly Thr Leu His Ala Gln His Val Ala Gly His Asp Ile Glu Ile Ile Asp Pro Thr Gly Ala Gly Asp Cys Phe Gly Ala Thr Phe Ile Thr Leu Phe Leu Ser Gly Phe Pro Ala His Lys Ala Leu Gln Tyr Ala Asn Ala Ser Gly Ala Leu Ala Val Met Arg Gln Gly Pro Met Glu Gly Ile Ser Ser Leu Ala Asp Ile Glu Asp Phe Leu Gln Gln His <210> <211> <212> PRT <213> Escherichia coli <400> Met Tyr Met Pro Gly Lys Gln Met Leu Cys Cys Ile Leu Ile Ser Ile Ile Ser Glu Gly Asp Met Lys Ile Phe Ile Ser Leu Phe Leu Phe Ile Ile Ser Thr Asn Ser Phe Ala Asp Asp Ile Thr His Ala Gly Val Val

Arg	Ile	Glu	Gly	Leu	Ile	Thr	Glu	Lys	Thr	Cys	Ile	Ile	Ser	Asp
Glu														_

50 55 60 5

Ser Lys Asn Phe Thr Val Asn Met Pro Asp Val Pro Ser Ser Val

Val

65 70 75 80

Arg Ser Ala Gly Asp Val Thr Glu Lys Val Tyr Phe Ser Ile Thr

Leu 85 90 95

Thr Arg Cys Gly Ser Asp Val Gly Asn Ala Tyr Ile Lys Phe Thr Gly

100 105 110

Asn Thr Val Ser Glu Asp Ala Ser Leu Tyr Lys Leu Glu Asp Gly Ser

· 115 12O 125

Val Glu Gly Leu Ala Leu Thr Ile Phe Asp Lys Asn Lys Gly Ser Ile

130 135 140

Ser Asn Asp Val Lys Ser Met Val Phe Ser Leu Thr Ser Ser Val
Asp
145
150
155

Asn Ile Leu His Phe Phe Ala Ala Tyr Lys Ala Leu Lys Asn Asn Val 40 165 170 175

Gln Pro Gly Asp Ala Asn Ala Ser Val Ser Phe Ile Val Thr Tyr Asp 45 180 185 190

10

<400> 15

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Ile Val Pro Leu Leu Val Trp Pro Thr Val Ile Ala Leu Ala Val Leu
10 20 25 30

Ile Phe Thr Leu Thr Phe Leu Ala Glu Ile Ile Phe Ser Phe Pro Leu 15 35 40 45

Leu Val Val Arg Ile Ser Leu Gln Glu Leu Gln Leu Glu Leu Val

20 50 55 60

Val Tyr Ala Leu Phe Phe Ser Val Met Gly Gly Ile Gly Trp Gln
Phe
25 65 70 75 80

Ser Arg Arg Thr Pro Pro Glu Leu Lys Asn Arg Leu His Cys Trp Leu 30 85 90 95

Val Phe Ser Pro Val Tyr Phe Trp Leu Ile Leu Ser Asn Phe Ile Leu 35 100 105 110

Tyr Ile Ser Pro Glu Lys Ser Ala Leu Leu Glu Asn Ile Arg Asn Phe
40 115 120 125

Phe Leu Thr Phe Val Trp Leu Pro Leu Asn Phe Ser Pro Phe Trp Pro
45 130 135 140

Gln Pro Trp Thr Asp Phe Val Gly Pro Ile Ser Ala Gln Leu Gly Phe
50 145 150 155

Ala Leu Gly Tyr Tyr Cys Gln Trp Arg Ser Lys Asn Arg Ser His Arg Lys Lys Trp Gly Asp Trp Val Thr Cys Leu Ser Leu Ala Ile Leu Ala Leu Gly Pro Leu Phe Asn Tyr Leu Gln <210> <211> <212> PRT <213> Escherichia coli <400> Met Lys Phe Asn Leu Ser Asn Leu Ser Ala Val Leu Leu Ala Ser Gly Met Leu Met Ser Thr Ala Val Thr Ala Ala Pro Gly Asp Ala Thr Gln Phe Gly Gly Ala Asp Thr Asp Trp Ser Thr Val Asp Tyr Pro Arg Leu Thr Asp Met Asp Asp Asn Val Asp Ser Met Gly Gly Lys Ile Arg Phe Thr Gly Arg Val Val Lys Ala Thr Cys Lys Val Ala Thr Asp Ser Lys Gln Ile Glu Val Val Leu Pro Val Val Pro Ser Asn Leu Phe Thr Gly

<400> 17

Ile Asp Val Glu Ala Gln Gly Ala Ser Asn Gln Thr Asp Phe Asn Ile 105 . 100 110 . 5 Asn Leu Thr Glu Cys Ser Asn Thr Asp Asp Gln Lys Ile Glu Phe 115 120 125 10 Phe Thr Gly Thr Ala Asp Ser Ala Asn Lys Thr Leu Ala Asn Glu Val 130 135 140 15 Glu Gly Ser Thr Asp Ala Asp Asn Ser Gly Asn Ala Gly Ala Thr Gly 145 150 155 - . 20 160 Val Gly Ile Arg Ile Tyr Ser Lys Gly Thr Thr Asn Asn Gly Leu Ile 25 165 170 175 Asn Leu Asn Thr Thr Ala Ala Glu Gly Ser Ala Ser Thr Ala Ala Tyr 30 180 185 190 Thr Ile Pro Gly Asn Ala Thr Thr His Asp Phe Ser Ala Ala Phe Thr 35 195 200 205 Ala Gly Tyr Ala Gln Asn Gly Ser Thr Val Ala Pro Gly Val Val Lys 40 210 215 220 Ser Thr Ala Ser Phe Val Val Leu Tyr Glu 225 230 45 <210> 17 <211> 336 <212> PRT50 <213> Escherichia coli

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		Pro	Ser	Asn	Trp	Ala	Ser	Gly	Thr	Glu	Gly	Tyr	Ile	Lys	Ile
5	Arg				165					170			ů,		175
	Ile Leu	Asp	Lys	Lys	Ile	Ile	Ser	Asp	Val	Ser	Leu	Ser	Asn	Val	Leu
10				180					185					190	
	Val Pro	Ser	Leu	Tyr	Val	Ser	Gln	Ile	Pro	Thr	Glu	His	Gly	Pro	Ile
15			195					200					205		
		Phe	Asn	Ala	Tyr	Ile	Gly	Asn	Leu	Asn	Ile	Gln	Val	Pro	Gln
20	Gly	210					215					220	• • • • • • • • • • • • • • • • • • • •		
	Cys Val 225	Thr	Ile	Asn	Glu	Gly 230	Thr	Ser	Phe	Thr	Val 235	Asn	Met	Pro	Asp
25	240					230					233				
	_	Ala	Ser	Glu	Leu	Ser	Arg	Ala	Gly	Ala	Gly	Ala	Lys	Pro	Ala
30	Gly		٠		245					250					255
	Val Asp	Thr	Pro	Val	Ala	Thr	Thr	Ile	Pro	Ile	Asn	Cys	Thr	Asn	Lys
35	1101			260					265					270	
	Thr Thr	Asp	Ala	Val	Met	Thr	Leu	Val	Phe	Asp	Gly	Asn	Ile	Ser	Ala
40			275					280				,	285		
	Arg Pro	Asp	Thr	Asn	Gly	Lys	Gln	Ser	Ile	Ile	Gln	Ala	Gln	Asp	Asn
45	110	29Ó					295			•		300			
	Asp	Val	Glv	Ile	Met	Ile	Met	Asp	Ser	Gln	Gln	Asn	Ser	Val	Asp

Asp Val Gly Ile Met Ile Met Asp Ser Gln Gln Asn Ser Val Asp Leu
50 305 320 310 310 315

	Asn Asn	Ala	Leu	Ala	Thr	Ser	Val	Gly	Val	Pro	Phe	Arg	Leu	Val	Glu	
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15	Lys 1				5					10					15	
		Met	Lys	Arg	Val	Val	Pro	Leu	Leu	Leu	Val	Ile	Met	Pro	Ala	
20	Cys	,		20					25					30		
٥٥		Ile	Ala	Gly	Met	Arg	Phe	Asn	Pro	Ala	Phe	Leu	Ser	Gly	Asp	
25	Thr		35				•	40			•		45			
		Ala	Val	Ala	Asp	Leu	Ser	Arg	Phe	Glu	Lys	Gly	Met	Thr	Tyr	
30	Leu	50					55					60				
2.5		Gly	Ser	Tyr	Glu	Val	Glu	: Val	Trp	Val	Asn	Asp	Ser	Pro	Leu	
35	Leu 65					70					75					80
		Arg	Thr	Val	Thr	Phe	Lys	Ala	Asp	Asp	Glu	Asn	Gln	Leu	Ile	
40	Pro				85					90					95	
15		Leu	Ser	Leu	Ala	Asp	Leu	Leu	Ser	Leu	Gly	Ile	Asn	Lys	Asn	
45	Ala			100					105			-		110		
	_	_	9 3	~ 7	~ J	-		_	~	~ 1	-m	~	~		_	

Leu Pro Glu Gln Ala Leu Ala Ser Ser Glu Asn Ser Cys Leu Asp

120

125

50

Leu

	_	Ile	Trp	Phe	Pro	Asp	Val	His	Tyr	Met	Pro	Glu	Leu	Asp	Ala
5	Gln	130				•	135					140			
	Arg Arg	Leu	Lys	Leu	Thr	Phe	Pro	Gln	Ala	Ile	Ile	Lys	Arg	Asp	Ala
10	145 160	,				150					155				
15	Gly Leu	Tyr	Ile	Pro	Pro	Glu	Gln	Trp	Asp	Asn	Gly	Ile	Thr	Ala	Phe
13	пец				165					170					175
20		Asn	Tyr	Asp	Phe	Ser	Gly	Asn	Asn	Asp	Arg	Gly	Asp	Tyr	Ser
20	Ser			180					185					190	-
25		Asn	Tyr	Tyr	Leu	Asn	Leu	Arg	Ala	Gly	Ile	Asn	Ile	Gly	Ala
25	Trp		195	٠				200					205		
20	_	Phe	Arg	Asp	Tyr	Ser	Thr	Trp	Ser	Arg	Gly	Ser	Asn	Ser	Ala
30	Gly	210			,		215					220			
25	-	Leu	Glu	His	Ile	Ser	Ser	Thr	Leu	Gln	Arg	Val	Ile	Ile	Pro
35	Phe 225 240		•			230					235				
40	_	Ser	Glu	Leu	Thr	Leu	Gly	Asp	Thr	Trp	Ser	Ser	Ser	Asp	Val
	Phe				245					250					255
45		Ser	Val	Ser	Ile	Arg	Gly	Ile	Lys	Leu	Glu	Ser	Asp	Glu	Asn
	Met			260	ı		•		265					270	
50	Leu Ala	Pro	Asp	Ser	Gln	Ser	Gly	Phe	Ala	Pro	Thr	Val	Arg	Gly	Ile

275	280	285

									•				•		
5	Lys Tyr	Ser	Arg	Ala	Gln	Val	Thr	Ile	Lys	Gln	Asn	Gly	Tyr	Val	Ile
	- 3 -	290					295					300			
10	Gln Pro	Thr	Tyr	Met	Pro	Pro	Gly	Pro	Phe	Glu	Ile	Ser	Asp	Leu	Asn
	305 320					310					315				
15	Thr Asn	Ser	Ser	Ala	Gly	Asp	Leu	Glu	Val	Thr	Ile	Lys	Glu	Ser	Asp
					325		,			330					335
20	Ser Gln	Glu	Thr	Val	Tyr	Thr	Val	Pro	Tyr	Aļa	Ala	Val	Pro	Ile	Leu
		٠		340					345					350	
25		Glu	Gly	His	Leu	Lys	Tyr	Ser	Thr	Thr	Val	Gly	Gln	Tyr	Arg
	Ser		355					360					365		
30	Asn Ile	Ser	Tyr	Asn	Gln	Lys	Ser	Pro	Tyr	Val	Phe	Gln	Gly	Glu	Leu
		370			•		375					380			
35	Trp Ser	Gly	Leu	Pro	Trp	Asp	Ile	Thr	Ala	Tyr	Gly	Gly	Ala	Gln	Phe
	385 400					390					395			٠	
40		Asp	Tyr	Arg	Ala	Leu	Ala	Leu	Gly	Leu	.Gly	Leu	Asn	Leu	Gly
	Val				405					410					415

Phe Gly Ala Thr Ser Phe Asp Val Thr Gln Ala Asn Ser Ser Leu Val

420
430

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	V	VO 200)5/0978	323				52/	370				PCT	/EP200	5/00397
	_	Gly	Ser	Lys	His	Gln	Gly	Gln	Ser	Tyr	Arg	Phe	Leu	Tyr	Ser
	Lys		435					440					445		
5	-		7	63	 .		57. 1	·n 7	1	** 1	T	** 7		-	_
	Ser Tyr		Val	GIn	Thr	GТĀ		Ala	Phe	His	Ile		GTĀ	Tyr	Arg
		450					455					460	•		
10	Ser	Thr	Gln	Gly	Phe	Tyr	Thr	Leu	Ser	Asp	Thr	Thr	Tyr	Gln	Gln
	Met 465					470					475				
15	480												٠		
	Ser	Gly	Thr	Val	Val	Asp	Pro	Lys	Thr	Leu	Asp	Asp	Lys	Asp	Tyr
	Val		•		485					490					495
20											•				
	Tyr Lys	Asn	Trp	Asn	Asp	Phe	Tyr	Asn	Leu	Arg	Tyr	Ser	Lys	Arg	Gly
25	_			500					505					510	
	Phe	Gln	Ala	Ser	Val	Ser	Gln	Pro	Phe	Gly	Asn	Tyr	Gly	Ser	Met
	Tyr		515					520					525		•
30															
	Leu Ser	Ser	Ala	Ser	Gln	Gln	Thr	Tyr	Trp	Asn	Thr	Asp	Lys	Lys	Asp
35		530					535					540			
	Leu	Tvr	Gln	Val	Glv	Tvr	Asn	Thr	Ser	Tle	Lys	Glv	Tle	Tvr	T _i eii
	Asn 545	-1-			241	550			201		555			- 1 -	10u,
40	560					550					333				
	T = T	73.7 ==	Ψrn	Agn	ጥኒንድ	Sar	T.v.c	Sar	Pro	Glv	Thr	7) en	7\] =	7) en	Twe
15	Ile	nia	115	ASII		Der	пуз	Der	110		7.117	ASII	ALG	ASP	
45					565					570					575
	Val Thr	Ser	Leu	Asn	Val	Ser	Leu	Pro	Ile	Ser	Asn	Trp	Leu	Ser	Ser

Thr

	Asn	Asp	Gly	Arg	Ser	Ser	Ser	Asn	Ala	Met	Thr	Ala	Thr	Туr	Glv
	Tyr		595	_				600	*				605		4
5															
	Ser Gly	Gln	Asp	Asn	His	Gly	Gln	Val	Asn	Gln	Tyr	Thr	Gly	Val	Ser
10		610					615					620			
•	Ser Phe	Leu	Leu	Glu	Gln	His	Asn	Leu	Ser	Tyr	Asn	Ile	Gln	His	Gly
15	625 640					630					635				
	•											•			
	Ala Arg	Asn	Gln	Asp	Asn	Ser	Ser	Ser	Gly	Ser	Val	Gly	Val	Asn	Tyr
20					645					650					655
	Gly Gly		Tyr	Gly	Ser	Leu	Asn	Ser	Ala	Tyr	Ser	Tyr	Asp	Asn	Glu
25	GIY			660					665					670	
	Asn	Gln	Gĺn	Tle	Asn	Tvr	Glv	Tle	Ser	Glv	ΔΊа	T.A.1	Vəl	Val	Uic
30	Glu	0111	675	220	21011	- y -	CLY	680	DCI	O _T y	1114	шси	685	vai	·
			0.0					000					005		
	Asn Ile	Gly	Leu	Thr	Leu	Ser	Gln	Pro	Leu	Gly	Glu	Thr	Asn	Val	Leu
35		690					695					700			•
	Lys	Ala	Pro	Gly	Ala	Asn	Asn	Val	Asp	Val	Gln	Ara	Glv	Thr	Glv
40	Ile 705			_		710			*		715	٦	4		4
	720														
	Ser	Thr	Asp	Trp	Arg	Gly	Tyr	Ala	Val	Val	Pro	Tyr	Ala	Thr	Glu
45	Tyr				725					730		-			735
				•											
50	Arg Thr	Arg	Asn	Asn ,	Ile	Ser	Leu	Asp	Pro	Met	Ser	Met	Asn	Met	His
,				740					745					750	

	Glu Leu	Leu	Asp	Ile	Thr	Ser	Thr	Glu	Val	Ile	Pro	Gly	Lys	Gly	Ala
5	Lou		755					760					765		
	Val	Arg	Ala	Glu	Phe	Ala	Ala	His	Ile	Glv	Ile	Ara	Glv	Ten	Phe
10	Thr	770					775			1		780	<u></u>	200	1110
		•													
	Val Ala	Arg	Tyr	Arg	Asn	Lys	Ser	Val	Pro	Phe	Gly	Ala	Thr	Ala	Ser _.
15	785 800					790					795				
						-									
20	Gln Gly	Ile	Lys	Asn	Ser	Ser	Gln	Ile	Thr	Gly	Ile	Val	Gly	Asp	Asn
	_				805					810					815
	Gln	Leu	Tyr	Leu	Ser	Gly	Leu	Pro	Leu	Glu	Gly	Val	Ile	Asn	Ile
25	Gln			820					825					830	
								•							
30	Trp Pro	Gly	Asp	Gly	Val	Gln	Gln	Lys	Cys	Gln	Ala	Asn	Tyr	Lys	Leu
			835					840					845		
	Glu	Thr	Glu	Leu	Asp	Asn	Pro	Val	Ser	Tyr	Ala	Thr	Leu	Glu.	Cys
35	Arg	850					855					860			
40	<210 19)> 1	L9 <2	211>	169	∂ <21	.2>	PRT	<213	3> E	Esche	erich	nia d	coli	<400>
		Gly	Ala	Ile	Tyr	Val	Lys	Arg	Leu	Ile	Leu	Ser	Val	Ala	Leu
	Ile 1				5					10					15
45		_		7.7	~	_		~	_		_			_	
	Ile Ser	Pro	TTE	Ala	ser	Asn	Ala	ser		ALa	Leu	Asn	Gln		Ser
50	•			20					25					30	

	W	O 2005	5/09782	23				55/37	70				PCT/	EP200	5/003972
	Ser Met	Leu	Asn	Asp	Gly	Val	Glu	Thr	Phe	Phe	Ile	Ser	Cys	Phe	Asp
	Met		35					40					45 .		
5	_												•		

Pro Gln Glu Thr Thr Asp Met Asp Ala Cys Gln Arg Val Gln Leu 50 55 60

10
Ala Gln Val Ser Trp Val Lys Asn Lys Tyr Ser Val Ala Ala Leu
Asn
65
70
75
80

Arg Leu Lys Gln Asp Asn Lys Asp Asp Pro Gln Arg Leu Gln Glu Leu

85
90
95

20
Thr Ala Ser Phe Asn Ala Glu Ser Glu Ala Trp Thr Glu Leu Ile Glu
100
105
110

Lys Ala Ser Lys Ser Val Gln Val Asp Tyr Val Gly Gly Thr Ile Ala 115 120 125

30
Gly Thr Ala Val Ala Ser Arg Gln Ile Gly Leu Leu Glu Leu Gln Ser
130
135
140

His Asp Ile Trp Glu His Trp Leu Arg Ser Arg Gly Leu Asn Ser Ser 145 150 155

Ser Phe Ala Arg Thr Lys Val Gln Ile 165

	Met Leu 1	Ala	Met	Phe	Thr 5	Pro	Ser	Phe	Ser	Gly 10	Leu	Lys	Gly	Arg	Ala 15	
5	_				Ü					10		•			10	
	Phe Val	Ser	Leu	Leu	Phe	Ala	Ala	Pro	Met	Ile	His	Ala	Thr	Asp	Ser	
				20					25					30		
10	Thr Thr	Thr	Lys	Asp	Gly	Glu	Thr	Ile	Thr	Val	Thr	Ala	Asp	Ala	Asn	
			35					40					45			
15 ⁻		Thr	Glu	Ala	Thr	Asp	Gly	Tyr	Gln	Pro	Leu	Ser	Thr	Ser	Thr	
	Ala	50					55					60			al ·	
20	Thr Thr	Leu	Thr	Asp	Met	Pro	Met	Leu	Asp	Ile	Pro	Gln	Val	Val	Asn	•
	65					70			٠		75					80
25	Val	Ser	Asp	Gln	Val	Leu	Glu	Asn	Gln	Asn	Ala	Thr	Thr	T.e.11	Asn	-
	Glu				85				0	90				200	95	
30																
	Ala Gly	Leu	Tyr	Asn	Val	Ser	Asn	Val	Val	Gln	Thr	Asn	Thr	Leu	Gly	ı
	4			100					105					110		
35	Thr	Gln	Asp	Ala	Phe	Val	Arg	Arg	Gly	Phe	Gly	Ala	Asn	Arq	Asp	
	Gly		115				•	120	_		_		125	,	2	
40																
	Ser Asn	Ile	Met	Thr	Asn	Gly	Leu	Arg	Thr	Val	Leu	Pro	Arg	Ser	Phe ·	
		130					135					140				
45	Ala Leu	Ala	Thr	Glu	Arg	Val	Glu	Val	Leu	Lys	Gly	Pro	Ala	Ser	Thr	
50	145 160					150					155					

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								57/3	70							
	Tyr Arg	Gly	Ile	Leu	Asp	Pro	Gly	Gly	Leu	Ile	Asn	Val	Val	Thr	Lys	
					165					170					175	
5		Glu	Lys	Thr	Phe	His	Gly	Ser	Val	Ser	Ala	Thr	Ser	Ser	Ser	
	Phe			180					185					190		
10		Gly	Gly	Thr	Gly	Gln	Leu	Asp	Ile	Thr	Gly	Pro	Ile	Glu	Gly	
	Thr		195					200				-	205			
15	Gln	Leu	Ala	Tyr	Arq	Leu	Thr	Glv	Glu	Val	Gln	Asp	Glu	Asp	Tvr	
	Trp	210		4	3		215	.				220			- , -	
20	7)	7)	Db a	<i>C</i> 1	T	<i>C</i> 3	7)	C	m la aa	'D1	T.	7 . 7 -	T.	9	T	•
	Thr 225	ASII	Pne	GTĀ	туз	Glu 230	Arg	ser	inr	Pne	235	Ala	Pro	Ser	ьeu	
25	240										200					
		Phe	Gly	Asp	Asn	Ala	Thr	Val	Thr	Met	Leu	Tyr	Ser	His	Arg	
30	Asp				245					250					255	
	Tyr	Lys	Thr	Pro	Phe	Asp	Arg	Gly	Thr	Ile	Phe	Asp	Leu	Thr	Thr	
35	Lys			260					265					270		
33	Gln	Pro	Val	Asn	Val	Asp	Arq	Lys	Ile	Ara	Phe	Asp	Glu	Pro	Phe	
	Asn		275				3	280					285			
40	- -1	m1	70			~	70		~ ·		_	_				
	His	290	Asp	GΤĀ	GIn	Ser	295	Leu	ALa	GIn	Leu	300	Ala	GLu	Tyr	
45		<u></u> V					2,70					,				
	Asp	Asn	Ser	Gln	Trp	Thr	Ala	Arg	Phe	Asp		Ser	Tyr	Ser	Gln	
50	305 320					310					315					

	Lys Thr	Tyr	Ser	Asp	Asn	Gln	Ala	Arg	Val	Thr	Ala	Tyr	Asp	Ala	Thr
5	7.117				325					330					335
		Thr	Leu	Thr	Arg	Arg	Val	Asp	Ala	Thr	Gln	Gly	Ser	Thr	Gln
10	Arg			340					345					350	
		His	Ala	Thr	Arg	Ala	Asp	Leu	Gln	Ġly	Asn	Val	Asp	Ile	Ala
15	Gly		355					360					365		
	Phe Leu	Tyr	Asn	Glu	Ile	Leu	Gly	Gly	Val	Ser	Tyr	Glu	Tyr	Tyr	Asp
20	шeu	370					375					380			
	Leu Ile	Arg	Thr	Asp	Met	Ile	Arg	Cys	Lys	Lys	Ala	Lys	Asp	Phe	Asn
25	385 400					390		•			395				
	ጥኒኒሎ	Nan	Dro	Val	Тиг	Clv	7\ c.n	Thr	S0 %	Tvo	Cvc	Thr	Thy	T - 77	C o 20
30	Ala	71311	110	vai	405	GIY	ASII	1147	Der	410	Суб	1111.	T11T	vaı	415
	G	7	C	71	Q 3	m1.	- 7		C 1		70		~		_
35	Ala	Asp	ser	420	GTU	rnr	тте	Lys	425	GIU	Asn	Tyr	Ser	A1a 430	Tyr
			,												
40	Gln Ile	Asp	Ala 435	Leu	Tyr	Leu	Thr	Asp 440	Asn	Trp	Ile	Ala	Val 445	Ala	Gly
10								440	•				110		
45	Arg Asn	Tyr 450	Gln	Tyr	Tyr	Thr	Gln 455	Tyr	Ala	Gly	Lys	Gly 460	Arg	Pro	Phe
13															
50	Leu	Asn	Thr	Asp	Ser		Asp	Glu	Gln	Trp		Pro	Lys	Leu	Gly
50	465 480					470					475				

	Val Gln	Tyr	Lys	Leu	Thr	Pro	Ser	Val	Ser	Leu	Phe	Ala	Asn	Tyr	Ser
5	GIII				485					490					495
	Thr	Phe	Met	Pro	Gln	Ser	Ser	Tle	Δla	Ser	ጥህዮ	Tle	Gly	Asn	, T. 211
	Pro	1110	1100	110	02.11	001	501	110	111 a	DCI	T	110	Оту	АЗР	леα
10				500					505	,				510	
		Glu	Ser	Ser	Asn	Ala	Tyr	Glu	Val	Gly	Ala	Lys	Phe	Glu	Leu
15	Phe		515					520					525	•	
		.													
		Gly	Ile	Thr	Ala	Asp	Ile	Ala	Leu	Phe	Asp	Ile	His	Lys	Arg
20	Asn	530					535					540			
															•
		Leu	Tyr	Thr	Glu	Ser	Ile	Gly	Asp	Glu	Thr	Ile	Ala	Lys	Thr
25	Ala 545					550					555				
	560														
	Glv	Ara	Val	Ara	Ser	Ara	Glv	Va l	Glu	Val	Asn	T.e.11	Ala	Glv	Z .] =
30	Leu	****9	Val	**** 9		1119	O L y	Val	OIG		1100	пса	ZII	Gry	
					565					570					575
	Thr	Glu	Asn	Ile	Asn	Ile	Ile	Ala	Ser	Tyr	Gly	Tyr	Thr	Asp	Ala
35	Lys									•	_	-		_	
				580					585					590	
	Val	Leu	Glu	Asp	Pro	Asp	Tyr	Ala	Gly	Lys	Pro	Leu	Pro	Asn	Val
40	Pro		595					600					605		
. ~		His	Thr	Gly	Ser	Leu	Phe	Leu	Thr	Tyr	Asp	Ile	His	Asn	Met
45	Pro	610					615					620			
50	Gly Arg	Asn	Asn	Thr	Leu	Thr	Phe	Gly	Gly	Gly	Gly	His	Gly	Val	Ser

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60/370 Arg Ser Ala Thr Asn Gly Ala Asp Tyr Tyr Leu Pro Gly Tyr Phe Val Ala Asp Ala Phe Ala Ala Tyr Lys Met Lys Leu Gln Tyr Pro Val Thr Leu Gln Leu Asn Val Lys Asn Leu Phe Asp Lys Thr Tyr Tyr Thr Ser Ser Ile Ala Thr Asn Asn Leu Gly Asn Gln Ile Gly Asp Pro Arg Glu Val Gln Phe Thr Val Lys Met Glu Phe 21 <211> 606 <212> PRT <213> Escherichia coli <400> <210> Met Lys Ile Ser Trp Asn Tyr Ile Phe Lys Asn Lys Trp Arg Phe His Ile Thr Ser Ile Ser Leu Phe Leu Ile Met Leu Ala Val Ser Ile Ala Phe Leu His Leu Arg Phe Asn Thr Leu Ser Ser Thr Asp Lys Met Arg Leu Glu Met Tyr Lys Ser Thr Leu Tyr Ser Thr Ile Glu Gln Phe

Tyr

Val Leu Pro Tyr Met Leu Ser Thr Asp His Ile Ile Arg Gln Ala Val

Ile Thr Pro Asp Asp Met Thr Ser Ser Glu Leu Asn Gln Arg Ile Ala

His Phe Asn Thr Gln Leu Lys Thr Ala Ala Ile Phe Ile Leu Asp Thr

Gln Gly Lys Ala Ile Ala Ser Ser Asn Trp Gln Asp Pro Gly Ser Tyr

Val Gly Gln Asn Tyr Ser Tyr Arg Pro Tyr Tyr Lys His Ala Met Ser

Gly Leu Asn Gly Arg Phe Tyr Gly Ile Gly Ser Thr Thr Asn Thr Pro

Gly Phe Phe Leu Ser Thr Ser Ile Lys Asp Lys Gly Lys Ile Val Gly

Val Val Val Lys Ile Ser Leu Asn Glu Ile Glu Lys Ala Trp Ala

Glu Gly Pro Glu Asn Ile Ile Val Asn Asp Glu His Gly Ile Ile Phe

Leu Ser Ser Lys Ser Pro Trp Arg Met Arg Thr Leu Gln Pro Leu Pro

	V	VO 200	5/0978	23		62/370							PCT/EP2005/0039			
5	Val Asp 225 240	Gln	Ala	Lys	Gln	Lys 230	Leu	Gln	Ser	Thr	Arg 235	Gln	T yr	Ser	Leu	
10	Asn Phe	Leu	Leu	Pro	Ala 245	Asp	Tyr	Tyr	Pro	Cys 250	Tyr	Thr	V.al	Ser	Asn 255	
15	Thr Tyr	Phe	Leu	Lys 260	Asp	Lys	Lys	Glu	Gln 265	Leu	Суз	Leu	Phe	Pro 270	Gln	
20	Tyr Ile	Thr	Gln 275	Gln	Ile	Ala	Ile	Pro 280	Glu	Phe	Asn	Trp	Lys 285	Met	Thr	
25	Met Val	Val 290	Pro	Leu	Asp	Asn	Leu 295	Tyr	Trp	Ser	Trp	Ala 300	I le	Ser	Leu	
30	Ile Trp 305 320	Thr	Leu	Ile	Ile	Tyr 310	Leu	Leu	Phe	Leu	Leu 315	Phe	Ile	Lys	Tyr	
35	Arg Thr	Met	Arg	Ser	His 325	Ala	Gln	Gln	Leu	Leu 330	Thr	Leu	Ala	Asn	Glu 335	
40	Leu Asn	Glu	Lys	Gln 340	Val	Lys	Glu	Arg	Thr 345	Ser	Ala	Leu	Glu	Leu 350		
45	Gln Val	Lys	Leu 355	Ile	Gln	Glu	Ile	Lys 360	Glu	Arg	Ser	Gln	Ala 365	Glu	Gln	

Leu Gln Ile Thr Arg Ser Glu Leu Ala Glu Ser Ser Lys Leu Ala Aļa 380 50 370 375

. 5	Leu Leu 385 400	Gly	Gln	Met	Ala	Thr 390	Glu	Ile	Ala	His	Glu 395	Gln	Asn	Gln	Pro
10	Ala Lys	Ala	Ile	His	Ala 405	Leu	Thr	Asp	Asn	Ala 410	Arg	Thr	Met	Leu	Lys 415
15	Glu Val	Met	Tyr	Pro 420	Gln	Val	Glu	Gln	Asn 425	Leu	Lys	His	Ile	Ile 430	Ser
20	Ile Ser	Glu	Arg 435	Met	Thr	Gln	Leu	Ile	Ser	Glu	Leu	Lys	Ala 445	Phe	Ala
25	Arg Tyr	His 450	Arg	Val	Pro		Gly 455	Ser	Ala	Asp	Val	Ile 460	Lys	Val	Met
30	Ser Glu 465 480	Ala	Val	Ala	Leu	Leu 470	Asn	His	Ser	Met	Glu 475	Lys	Asn	Asn	Ile
35	Arg Glu	Arg	Ile	Lys	Ala 485	Pro	Ser	Met	Pro	Leu 490	Phe	Val	Asn	Cys	Asp 495
40	Leu Asp	Gly	Leu	Glu 500	Gln	Ile	Phe	Ser	Asn 505	Leu	Ile	Ser	Asn	Ala 510	Leu
45	Ser Gln	Met	Glu 515	Gly	Ser	Ser	Tyr	Lys 520	Arg	Leu	Asp	Ile	Ala 525	Ile	Arg
50	Ala Phe	Asn 530	Asn	Lys	Val	Ile	Ile 535	Thr	Ile	Lys	Asp	Ser 540	Gly	Gly	Gly

5	Ala Lys 545 560	Pro	Glu	Val	Val	Asp 550	Arg	Ile	Phe	Glu	Pro 555	Phe	Phe	Thr	Thr	
10	Arg Arg	Arg	Gly	Met	Gly 565	Leu	Gly	Leu	Ala	Ile 570	Val	Ser	Glu	Ile	Val 575	
15	Asn Ala	Ser	Asn	Gly 580	Ala	Leu	His	Ala 、	Ser 585	Asn	His	Pro	Glu	Gly 590	Gly	
20	Val	Met	Thr 595	Leu	Thr	Trp	Pro	Glu 600	Trp	Gly	Glu	Glu	His 605	Glu		
25	<210 22	0> 2	22 <2	211>	103	L <2:	L2>	PRT	<213	3> I	Esche	erich	nia d	coli	. <40	00>
25	Val Leu 1	Leu	Thr	Pro	Gln 5	His	Leu	Arg	Cys	Val	Leu	Thr	Cys	Ser	Asp	
30										,						
	Leu Phe	Thr	Leu	Leu	Ser	Gly	Thr	Val	Met	Ser	Gln	Met	Pro	Leu	Tyr	
	rne			20					25					30		
35																
	Leu Ile	Asn	Thr	Gln	Lys	Lys	Leu	Thr	Ala	His	Tyr	Glu [,]	Trp	Leu	Gln	
			35					40					45			
40	7 ~ ~	T 011	mb z	7\ a.s.	Пр ж	M	C1	T 0	77- 1	T	7	Т	7.4 - L	D.		
	Pro		Thr	Asp	TIIT	т ў т		ьеи	Val	ту	Arg		Met	Pro	тте	
		50					55					60				
45		Leu	Asp	Val	Val	Val	Lys	Val	Gly	Lys	Leu	Val	Leu	Pro	Glu	
	Lys 65		i)			70					75					80

Gly His His Gly Phe Tyr Pro Glu Ala Gly Val Val Tyr Arg Thr Val

Ala Pro Glu Asn Pro

<210> 23 <211> 263 <212> PRT <213> Escherichia coli <400>

Met Met Lys Asn Thr Gly Tyr Ile Leu Ala Leu Cys Leu Thr Ala 1 .

Gly His Val Leu Ala His Asp Val Trp Ile Thr Gly Lys Gln Ala

Asn Asn Val Thr Ala Glu Ile Gly Tyr Gly His Asn Phe Pro Ser Lys

Gly Thr Ile Pro Asp Arg Arg Asp Phe Phe Glu Asn Pro Arg Leu Tyr

Asn Gly Lys Glu Thr Ile Thr Leu Lys Pro Ala Ser Thr Asp Tyr Val

Tyr Lys Thr Glu Ser Ala Ser Lys Asp Asn Gly Tyr Val Leu Ser Thr

Tyr Met Lys Pro Gly Tyr Trp Ser Arg Thr Ser Ser Gly Trp Lys Pro

Val Ser Arg Glu Gly Arg Asn Asp Val Ala Tyr Cys Glu Phe Val Thr

66/37

Lys Tyr Ala Lys Ser Phe Ile Pro Gly Glu Gln Gln Met Pro Ala Gln Leu Tyr Gln Ser Pro Thr Gly His Glu Leu Glu Ile Ile Pro Leu

Ser

Asp Ile Ser Arg Phe Ser Glu Asn Val Lys Leu Lys Val Leu Tyr Lys

Thr Ser Pro Leu Ala Gly Ala Ile Met Glu Leu Asp Ser Val Ser

Leu Thr Ser Ser Arg His Thr His Ala Val Glu His Lys His Pro Val

His Lys Ala Glu Leu Thr Phe Val Thr Asn Glu Asp Gly Ile Val

Val Pro Ser Leu His Ile Gly Gln Trp Leu Ala Lys Val Gln Asn Lys 230 .

Lys Ser Phe Gln Asp Lys Ser Leu Cys Asp Glu Thr Val Asp Val Ala

Thr Leu Ser Phe Ser Arg Asn

<210> 24 <211> 378 <212> PRT <213> Escherichia coli <400>

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	67/370	

Met Gly Lys Ile Lys Tyr Trp Leu Ile Val Gly Phe Ile Ile Leu Phe Ala Ile Phe Tyr Ile Ala Ile Ser Asp Arg Asp Ser Thr Leu Ser Arg Leu Lys Ser Ala Gly Glu Asn Gly Asp Val Glu Ala Gln Tyr Ala Leu Gly Leu Met Tyr Leu Tyr Gly Glu Ile Leu Asp Val Asp Tyr Gln Gln , 50. Ala Lys Ile Trp Tyr Glu Lys Ala Ala Asp Gln Asn Asp Pro Arg Ala Gln Ala Lys Leu Gly Val Met Tyr Ala Asn Gly Leu Gly Val Asn Gln Asp Tyr Gln Gln Ser Lys Leu Trp Tyr Glu Lys Ala Ala Gln Asn Asp Val Asp Ala Gln Phe Leu Leu Gly Glu Met Tyr Asp Asp Gly Leu Gly Val Ser Gln Asp Tyr Gln His Ala Lys Met Trp Tyr Glu Lys Ala Ala Ala Gln Asn Asp Glu Arg Ala Gln Val Asn Leu Ala Val Leu

Tyr

68/370

	Ala Trp	Lys	Gly	Asn	Gly	Val	Glu	Gln	Asp	Tyr	Arg	Gln	Ala	Γλε	Ser
					165					170					175
5	Tyr Leu	Glu	Lys	Ala	Ala	Ala	Gln	Asn	Ser	Pro	Asp	Ala	Gln	Phe	Ala
				180					185					190	
10	Gly Gln	Ile	Leu	Tyr	Ala	Asn	Ala	Asn	Gly	Val	Glu	Gln	Asp	Tyr	Gln
	0		195					200					205		
15	Ala	Lys	Asp	Trp	Tvr	Glu	Lvs	Ala	Ala	Glu	Gln	Asn	Phe	Ala	Asn
	Ala	210	1		- 1		215					220			11011
20	7														
	Gln Gln	Phe	Asn	Leu	Gly	Met	Leu	Tyr	Tyr	Lys	Gly	Glu	Gly	Val	Lys
	225 240					230					235				
25															
	Asn Asn	Phe	Arg	Gln	Ala	Arg	Glu	Trp	Phe	Glu	Lys	Ala	Ala	Ser	Gln
30					245					250					255
	Gln Gln	Pro	Asn	Ala	Gln	Tyr	Asn	Leu	Gly	Gln	Ile	Tyr	Tyr	Tyr	Gly
35				260					265					270	
	Gly	Val	Thr	Gln	Ser	Tyr	Arg	Gln	Ala	Lys	Asp	Trp	Phe	Glu	Lys
	Ala		275					280					285		
40		•												·	
	Ala Tyr	Glu.	Lys	Gly	His	Val	Asp	Ala	Gln	Tyr	Asn	Leu	Gly	Val	Ile
45		290					295					300		•	
	Glu Trp	Asn	Gly	Glu	Gly	Val	Ser	Gln	Asn	Tyr	Gln	Gln	Ala	Lys	Ala
50	305 320					310					315				

Tyr Glu Lys Ala Ala Ser Gln Asn Asp Ala Gln Ala Gln Phe Glu

Gly Val Met Asn Glu Leu Gly Gln Gly Glu Ser Ile Asp Leu Lys Gln

Ala Arg His Tyr Tyr Glu Arg Ser Cys Asn Asn Gly Leu Lys Lys

Cys Glu Arg Leu Lys Glu Leu Leu Tyr Lys

25 <211> 654 <212> PRT <213> Escherichia coli <400> <210>

Met Asn Val Ile Arg Thr Val Ile Cys Thr Leu Ile Ile Leu Pro Val

Gly Leu Gln Ala Ala Thr Ser His Ser Ser Met Val Lys Asp Thr Ile

Thr Ile Val Ala Thr Gly Asn Gln Asn Thr Val Phe Glu Thr Pro Ser

Val

Met Val Ser Val Val Thr Asn Asp Thr Pro Trp Ser Gln Asn Ala

Thr Ser Ala Gly Met Leu Lys Gly Val Ala Gly Leu Ser Gln Thr Gly

Ala Gly Arg Thr Asn Gly Gln Thr Phe Asn Leu Arg Gly Tyr Asp Lys

	Ser Met	Gly	Val	Leu	Val	Leu	Val	Asp	Gly	Val	Arg	Gln	Leu	Ser	Asp.	
5				100					105					110		
		Lys	Ser	Ser	Gly	Thr	Tyr	Leu	Asp	Pro	Ala	Leu	Val	Lys	Arg	
10	Ile	•	115					120					125			
	Glu	Val	Val	Arg	Gly	Pro	Asn	Ser	Ser	Leu	Tyr	Gly	Ser	Gly	Gly	
15	Leu	13,0					135					140				
	Gly	Gly	Val	Val	Asp	Phe	Ara	Thr	Ala	Asp	Ala	Ala	Asp	Phe	Leu	
20	Pro 145	-\$-			4	150					155		1101		104	
	. 160															
25	Pro Gly	Gly	Glu	Thr	Asn	Gİy	Leu	Ser	Leu	Trp	Gly	Asn	Ile	Ala	Ser	
	.				165	•				170					175	
30		His	Ser	Thr	Gly	Ser	Gly	Leu	Thr	Trp	Phe	Gly	Lys	Thr	GLy	
50	Lys			180					185		•			190		
		Asp	Ala	Leu	Leu	Ser	Val	Ile	Met	Arg	Lys	Arg	Gly	Asn	ILe	
35	Tyr		195	•				200					205			
	Gln	Ser	Asp	Gly	Glu	His	Ala	Pro	Asn	Lÿs	Glu	Lys	Pro	Ala	Ala	
40	Leu	210					215			-		220				
	Phe	Δla	T.ve	Glv	Sar	Val	Clu	Ile	ሞኮኮ	7.50	Sor	7) an	Tvo	7.7.	Cl	
45	Ala 225	пта	цуз		Der	230		116	TIIL	Asp	235	ASII	пур	ALA	GT À	
	240															
50	Ser	Leu	Arg	Leu	Tyr	Arg	Asn	Asn	Thr	Thr	Glu	Pro	Gly	Asn	Se r	

Thr

Gln Thr His Gly Asp Ser Gly Leu Arg Asp Arg Lys Thr Val Gln 5 Asn 260 265 270

Asp Val Gln Phe Trp Tyr Gln Tyr Ala Pro Val Asp Asn Ser Leu 10 lle $275 \hspace{1.5cm} 280 \hspace{1.5cm} 285$

Asn Val Lys Ser Thr Leu Tyr Leu Ser Asp Ile Thr Ile Lys Thr
15 Asn
290 295 300

Gly His Asn Lys Thr Ala Glu Trp Arg Asn Asn Arg Thr Ser Gly
Val
305 310 315

25 Asn Val Val Asn Arg Ser His Thr Leu Ile Phe Pro Gly Ala His Gln 325 330 335

30 Leu Ser Tyr Gly Ala Glu Tyr Tyr Arg Gln Gln Gln Lys Pro Glu Gly
340 345 350

35 Ser Ala Thr Leu Tyr Pro Glu Gly Asn Ile Asp Phe Thr Ser Leu Tyr 355 360 365

40 Phe Gln Asp Glu Met Thr Met Lys Ser Tyr Pro Val Asn Ile Ile Val 370 375 380

Glu Ser Arg Tyr Asp Arg Tyr Lys Ser Phe Asn Pro Arg Ala Gly Glu 385 390 395

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	Leu Pro	Lys	Ala	Glu		Leu	Ser	Pro	Arg	Ala	Ala	Ile	Ser	Val	Ser
_					405					410					415
5	Thr Ala	Asp	Trp	Leu	Met	Met	Tyr	Gly	Ser	Ile	Ser	Ser	Ala	Phe	Arg
	4			420					425					430	
10	Pro	Thr	Met	Ala	Glu	Met	Tyr	Arg	Asp	Asp	Val	His	Phe	Tyr	Arg
	Lys		435					440					445		
15	Gly	Lys	Pro	Asn	Tyr	Trp	Val	Pro	Asn	Leu	Asn	Leu	Lys	Pro	Glu
	Asn	450	•				455					460			
20	Asn	Tle	Thr	Ara	Glu	Tle	Glv	Δla	Glv	Tle	Gln	T,e11	Asn	Glv	T. 211
	Leu 465			9		470	O j	1120	CLY	110	475	100	1100	Ory	шси
25	480									•					
	Thr Asp	Asp	Asn	Asp	Arg	Leu	Gln	Leu	Lys	Gly	Gly	Tyr	Phe	Gly	Thr
30					485					490		•			495
	Ala Ser	Arg	Asn	Tyr	Ile	Ala	Thr	Arg	Val	Asp	Met	Lys	Arg	Met	Arg
35	ber			500					505					510	
		Ser	Tyr	Asn	Val	Ser	Arg	Ala	Arg	Ile	Trp	Gly	Trp	Asp	Met
40	Gln		515					520					525		
40	Gly	Asn	Tyr	Gln	Ser	Asp	Tyr	Val	Asp	Trp	Met	Leu	Ser	Tyr	Asn
45	Arg	530	-			-	535		-	-		540			
13	Thr	Glu	Ser	Met	Asp	Ala	Ser	Ser	Arg	Glu	Trp	Leu	Gly	Ser	Gly
50	Asn 545 560				_	550	٠	,	-	¥	555		-		-

VV C 2003/07/023		I C 1/E1 2003
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Pro Asp Thr Leu Ile Ser Asp Ile Ser Ile Pro Val Gly His Arq Gly

Val Tyr Ala Gly Trp Arg Ala Glu Leu Ser Ala Ser Ala Thr His Val

Lys Lys Gly Asp Pro His Gln Ala Gly Tyr Thr Ile His Ser Phe Ser

Leu Ser Tyr Lys Pro Val Ser Val Lys Gly Phe Glu Ala Ser Val Thr

Leu Asp Asn Ala Phe Asn Lys Leu Ala Met Asn Gly Lys Gly Val

> Leu Ser Gly Arg Thr Val Ser Leu Tyr Thr Arg Tyr Gln Trp

26 <211> 1376 <212> PRT <213> Escherichia coli <210> <400>

Met Asn Lys Ile Tyr Ala Leu Lys Tyr Cys Tyr Ile Thr Asn Thr Val

Lys Val Val Ser Glu Leu Ala Arg Arg Val Cys Lys Gly Ser Thr Arg

Arg Gly Lys Arg Leu Ser Val Leu Thr Ser Leu Ala Leu Ser Ala Leu

Leu Pro Thr Val Ala Gly Ala Ser Thr Val Gly Gly Asn Asn Pro Tyr

50 55 60

Gln Thr Tyr Arg Asp Phe Ala Glu Asn Lys Gly Gln Phe Gln Ala Gly Ala Thr Asn Ile Pro Ile Phe Asn Asn Lys Gly Glu Leu Val Gly His Leu Asp Lys Ala Pro Met Val Asp Phe Ser Ser Val Asn Val Ser Ser Asn Pro Gly Val Ala Thr Leu Ile Asn Pro Gln Tyr Ile Ala Ser Val Lys His Asn Lys Gly Tyr Gln Ser Val Ser Phe Gly Asp Gly Gln Asn Ser Tyr His Ile Val Asp Arg Asn Glu His Ser Ser Ser Asp Leu His Thr Pro Arg Leu Asp Lys Leu Val Thr Glu Val Ala Pro Ala Thr Val Thr Ser Ser Ser Thr Ala Asp Ile Leu Asn Pro Ser Lys Tyr Ser Ala 190. Phe Tyr Arg Ala Gly Ser Gly Ser Gln Tyr Ile Gln Asp Ser Gln Gly

Lys Arg His Trp Val Thr Gly Gly Tyr Gly Tyr Leu Thr Gly Gly

Ile.

210 215 220

Leu Pro Thr Ser Phe Phe Tyr His Gly Ser Asp Gly Ile Gln Leu
5 Tyr
225 230 235
240

- 10 Met Gly Gly Asn Ile His Asp His Ser Ile Leu Pro Ser Phe Gly Glu
 245 250 255
- 15 Ala Gly Asp Ser Gly Ser Pro Leu Phe Gly Trp Asn Thr Ala Lys Gly
 260 265 270
- 20 Gln Trp Glu Leu Val Gly Val Tyr Ser Gly Val Gly Gly Thr Asn 275 280 285
- 25 Leu Ile Tyr Ser Leu Ile Pro Gln Ser Phe Leu Ser Gln Ile Tyr Ser 290 295 300
- 30 Glu Asp Asn Asp Ala Pro Val Phe Phe Asn Ala Ser Ser Gly Ala Pro 305 310 315
- Leu Gln Trp Lys Phe Asp Ser Ser Thr Gly Thr Gly Ser Leu Lys Gln

 325
 330
 335
- Gly Ser Asp Glu Tyr Ala Met His Gly Gln Lys Gly Ser Asp Leu Asn
 340
 345
 350
- Ala Gly Lys Asn Leu Thr Phe Leu Gly His Asn Gly Gln Ile Asp Leu
 355 360 365

	V	VO 200	5/0978	23			,	76/3	370		PCT/EP2005/003972				
	Glu Asp	Asn	Ser	Val	Thr	Gln	Gly	Ala	Gly	Ser	Leu	Thr	Phe	Thr	Asp
	АЗР	370					375					380			
5	Tyr Ile 385 400	Thr	Val	Thr	Thr	Ser 390	Asn	Gly	Ser	Thr	Trp 395	Thr	Gly	Ala	Gly

Ile Val Asp Lys Asp Ala Ser Val Asn Trp Gln Val Asn Gly Val Lys

405
410
415

Gly Asp Asn Leu His Lys Ile Gly Glu Gly Thr Leu Val Val Gln Gly
420 425 430

Thr Gly Val Asn Glu Gly Gly Leu Lys Val Gly Asp Gly Thr Val Val 435 440 445

20

40

50

Leu Asn Gln Gln Ala Asp Ser Ser Gly His Val Gln Ala Phe Ser Ser 450 455 460

Val Asn Ile Ala Ser Gly Arg Pro Thr Val Val Leu Ala Asp Asn Gln
465 470 475
35 480

Gln Val Asn Pro Asp Asn Ile Ser Trp Gly Tyr Arg Gly Gly Val Leu 485 490 495

Asp Val Asn Gly Asn Asp Leu Thr Phe His Lys Leu Asn Ala Ala Asp 45 500 505 510

Tyr Gly Ala Thr Leu Gly Asn Ser Ser Asp Lys Thr Ala Asn Ile
Thr
515 520 525

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Leu Asp Tyr Gln Thr Arg Pro Ala Asp Val Lys Val Asn Glu Trp Ser Ser Ser Asn Arg Gly Thr Val Gly Ser Leu Tyr Ile Tyr Asn Asn Pro Tyr Thr His Thr Val Asp Tyr Phe Ile Leu Lys Thr Ser Ser Tyr Gly Trp Phe Pro Thr Gly Gln Val Ser Asn Glu His Trp Glu Tyr Val Gly His Asp Gln Asn Ser Ala Gln Ala Leu Leu Ala Asn Arg Ile Asn Asn Lys Gly Tyr Leu Tyr His Gly Lys Leu Leu Gly Asn Ile Asn Phe Ser Asn Lys Ala Thr Pro Gly Thr Thr Gly Ala Leu Val Met Asp Gly Ala Asn Met Ser Gly Thr Phe Thr Gln Glu Asn Gly Arg Leu Thr Ile Gln Gly His Pro Val Ile His Ala Ser Thr Ser Gln Ser Ile Ala Asn Thr Val Ser Ser Leu Gly Asp Asn Ser Val Leu Thr Gln Pro Thr Ser

	Phe Leu	Thr	Gln	Asp	Asp	Trp	Glu	Asn	Arg	Thr	Phe	Ser	Phe	Gly	Ser
5		690					695					700			
	Val Asn	Leu	Lys	Asp	Thr	Asp	Phe	Gly	Leu	Gly	Arg	Asn	Ala	Thr	Leu
10	705 720					710					715				
15	Thr Arg	Thr	Ile	Gln	Ala	Asp	Asn	Ser	Ser	Val	Thr	Leu	Gly	Asp	Ser
	٥				725					730			4		735
20	Val Glu	Phe	Ile	Asp	Lys	Lys	Asp	Gly	Gln	Gly	Thr	Ala	Phe	Thr	Leu
				740					745					750	
25	Glu Asn	Gly	Thr	Ser	Val	Ala	Thr	Lys	Asp	Ala	Asp	Lys	Ser	Val	Phe
	11011		755					760					765		
30	Gly Ile	Thr	Val	Asn	Leu	Asp	Asn	Gln	Ser	Val	Leu	Asn	Ile	Asn	Glu
50		770					775					780			
35	Phe Ser	Asn	Gly	Gly	Ile	Gln	Ala	Asn	Asn	Ser	Thr	Val	Asn	Ile	Ser
33	785 800					790					795				
40	Asp Asn	Ser	Ala	Val	Leu	Glu	Asn	Ser	Thr	Leu	Thr	Ser	Thr	Ala	Leu
					805					810					815
45	Leu Asp		Lys	Gly	Ala	Asn	Val	Leu	Ala	Ser	Gln	Ser	Phe	Val	Ser
	110P	•		820					825			,		830	
50	Gly Pro	Pro	Val	Asn	Ile	Ser	Asp	Ala	Thr	Leu	Ser	Leu	Asn	Ser	Arg

835	8 4 0	845

Asp Glu Val Ser His Thr Leu Leu Pro Val Tyr Asp Tyr Ala Gly
5 Ser 850 855 860

Trp Asn Leu Lys Gly Asp Asp Ala Arg Leu Asn Val Gly Pro Tyr

Ser

865
870
880

15 Met Leu Ser Gly Asn Ile Asn Val Gln Asp Lys Gly Thr Val Thr Leu 885 890 895

20 Gly Gly Glu Gly Glu Leu Ser Pro Asp Leu Thr Leu Gln Asn Gln Met
900 905 910

25 Leu Tyr Ser Leu Phe Asn Gly Tyr Arg Asn Thr Trp Ser Gly Ser Leu
915 920 925

30 Asn Ala Pro Asp Ala Thr Val Ser Met Thr Asp Thr Gln Trp Ser Met 930 935 940

 C_{i}^{f}

Asn Gly Asn Ser Thr Ala Gly Asn Met Lys Leu Asn Arg Thr Ile
Val
945 950 955
960

Gly Phe Asn Gly Gly Thr Ser Ser Phe Thr Thr Leu Thr Thr Asp Asn
965 970 975

Leu Asp Ala Val Gln Ser Ala Phe Val Met Arg Thr Asp Leu Asn Lys

980

985

990

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Ala Asp Lys Leu Val Ile Asn Lys Ser Ala Thr Gly His Asp Asn

Ser 995 1000 1005

Ile Trp Val Asn Phe Leu Lys Lys Pro Ser Asp Lys Asp Thr Leu 1010 1015 1020

10 Asp Ile Pro Leu Val Ser Ala Pro Glu Ala Thr Ala Asp Asn Leu 1025 1030 1035

Phe Arg Ala Ser Thr Arg Val Val Gly Phe Ser Asp Val Thr Pro 15 1040 1045 1050

Thr Leu Ser Val Arg Lys Glu Asp Gly Lys Lys Glu Trp Val Leu 1055 1060 1065

20

40

45

.

Asp Gly Tyr Gln Val Ala Arg Asn Asp Gly Gln Gly Lys Ala Ala 1070 1075 1080

25
Ala Thr Phe Met His Ile Ser Tyr Asn Asn Phe Ile Thr Glu Val
1085
1090
1095

30 Asn Asn Leu Asn Lys Arg Met Gly Asp Leu Arg Asp Ile Asn Gly 1100 1105 1110

Glu Ala Gly Thr Trp Val Arg Leu Leu Asn Gly Ser Gly Ser Ala 35 1115 1120 1125

Asp Gly Gly Phe Thr Asp His Tyr Thr Leu Leu Gln Met Gly Ala 1130 1135 1140

Asp Arg Lys His Glu Leu Gly Ser Met Asp Leu Phe Thr Gly Val 1145 1150 1155

Met Ala Thr Tyr Thr Asp Thr Asp Ala Ser Ala Gly Leu Tyr Ser 1160 1165 1170

50 Gly Lys Thr Lys Ser Trp Gly Gly Gly Phe Tyr Ala Ser Gly Leu 1175 1180 1185

5	Phe	Arg 1190		Gly	Ala	Tyr	Phe 1195	Asp	Leu	Ile	Ala	Lys 1200	Tyr	Ile	His
3	Asn	Glu 1205		Lys	Туг	Asp	Leu 1210	Asn	Phe	Ala	Gly	Ala 1215	Gly	Lys	Gln
10	Asn	Phe 1220	Arg	Ser	His	Ser	Leu 1225	Tyr	Ala	Gly	Ala	Glu 1230	Val	Gly	Tyr
15	Arg	Tyr 1235	His	Leu	Thr	Asp	Thr 1240		Phe	Val	Glu	Pro 1245	Gln	Ala	Glu
20	Leu	Val 1250	Trp	Gly	Arg	Leu	Gln 1255		Gln	Thr		Asn 1260	Trp	Asn	Asp
. 25	Ser	Gly 1265	Met	Asp	Val	Ser	Met 1270	Arg	Arg	Asn	Ser	Val 1275	Asn	Pro	Leu
	Val	Gly 1280	Arg	Thr	Gly	Val	Val 1285		Gly	Lys	Thr	Phe 1290	Ser	Gly	Lys
30	Asp	Trp 1295	Ser	Leu	Thr	Ala	Arg 1300	Ala	Gly	Leu	His	Tyr 1305	Glu	Phe	Asp
35	Leu	Thr 1310	Asp	Ser	Ala	Asp	Val 1315	His	Leu	Lys	Asp	Ala 1320	Ala	Gly	Glu
40	His	Gln 1325	Ile	Asn	Gly	Arg	Lys 1330	Asp	Gly	Arg	Met	Leu 1335	Tyr	Gly	Val
45	Gly	Leu 1340	Asn	Ala	Arg	Phe	Gly 1345	Asp	Asn	Thr	Arg	Leu 1350	Gly	Leu	Glu
	Val	Glu 1355	Arg	Ser	Ala	Phe	Gly 1360		Tyr	Asn	Thr	Asp 1365		Ala	Ile
50	Asn	Ala	Asn	Ile	Arg	Tyr	Ser	Phe							•

02/3/0

<210> 27 <211> 349 <212> PRT <213> Escherichia coli <400> Met Ile Thr Leu Phe Arg Leu Leu Ala Ile Leu Cys Leu Phe Phe Val Ser Ala Phe Ala Val Asp Cys Tyr Gln Asp Gly Tyr Arg Gly Thr Leu Ile Asn Gly Asp Leu Pro Thr Phe Lys Ile Pro Glu Asn Ala Gln Pro Gly Gln Lys Ile Trp Glu Ser Gly Asp Ile Asn Ile Thr Val Tyr Cys Asp Asn Ala Pro Gly Trp Ser Ser Asn Asn Pro Ser Glu Asn Val Tyr Ala Trp Ile Lys Leu Pro Gln Ile Asn Ser Ala Asp Met Leu Asn Asn Pro Tyr Leu Thr Phe Gly Val Thr Tyr Asn Gly Val Asp Tyr

Glu Gly Thr Asn Glu Lys Ile Asp Thr His Ala Cys Leu Asp Lys Tyr 115 . 120 125

Glu Gln Tyr Tyr Asn Gly Tyr Tyr His Asp Pro Val Cys Asn Gly Ser 130 135

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								03/3/	, 0						
	Val		Gln	Lys	Asn		Thr	Phe	Asn	Ala		Phe	Arg	Val	Tyr
5	145 160					150					155				
	Lys Thr	Phe	Lys	Ser		Pro	Ala	Gly	Asp	Gln	Thr	Val	Asn	Phe	Gly
10					165			,		170					175
-	Val Pro	Asn	Val	Leu	Gln	Phe	Asp	Gly	Glu	Gly	Gly	Ala	Asn	Met	Ala
15		,		180					185					190	
	Asn Ser	Ala	Lys	Asn	Leu	Arg	Tyr	Ala	Ile	Thr	Gly	Leu	Asp	Asn	Ile
20			195					200					205		
	Phe Ile	Leu	Asp	Cys	Ser	Val	Asp	Val	Arg	Ile	Ser	Pro	Glu	Ser	Gln
25		210					215					220			
	Val Pro	Asn	Phe	Gly	Gln	Ile	Ala	Ala	Asn	Ser	Ile	Ala	Thr	Phe	Pro
30	225 240			•• •		230					235				
	Lys Cys	Ala	Ala	Phe	Ser	Val	Ser	Thr	Ile	Lys	Asp	Ile	Ala	Ser	Asp
35	CYS				245					250		,			255
	Thr Leu	Glu	Gln	Phe	Asp	Val	Ala	Thr	Ser	Phe	Phe	Thr	Ser	Asp	Thr
40				260					265					270	
	Tyr Ile	Asp	Asn	Thr	His	Leu	Glu	Ile	Gly	Asn	Gly	Leu	Leu	Met	Arg
45			275					280					285		
	Thr	Asp	Gln	Lys	Thr	Gln	Glu	Asp	Ile	Lys	Phe	Asn	Gln	Phe	Lys
50	Leu	290					295					300			

	Asp	Ser	Thr	Tyr	Ile	Pro	Gly	Gln	Ser	Ala		Met	Ala	Thr	Arg	
5	305 320					310	-	,			315					
	Tyr Gly	Gln	Ala	Glu	Leu	Thr	Gln	Lys	Pro	Gly	Glu	Pro	Leu	Val	Tyr	
10					325					3,30					335	
15		Phe	Gln	Lys 340	Asp	Leu	Ile	Val	Lys 345	Ile	Asn	Tyr	Hịs			
	<210 28)> 2	28 <2	211>	840) <23	L2>	PRT	<213	3> I	Esch	erich	nia d	coli	< 40	00>
20	Met Ile	Asn	Asn	Lys	Asn	Thr	Phe	Ser	Arg	Asp	Lys	Leu	Ser	His	Ala	
v	1				5					10					15	
25	Lys Pro	Asn	Ala	Leu	Ser	Gly	Val	Val	Cys	Ser	Leu	Leu	Phe	Val	Leu	
				20					25					30		
30	Val Arg	His	Ala	Val	Glu	Phe	Asn	Val	Asp	Met	Ile	Asp	Ala	Glu	Asp	
	2	•	35					40					45			
35 .	Glu Pro	Asn	Ile	Asp	Ile	Ser	Arg	Phe	Glu	Lys	Lys	Gly	Tyr	Ile	Pro	
٠		50					55					60				
40	Gly Gln	Arg	Tyr	Leu	Val	Arg	Val	Gln	Ile	Asn	Lys	Asn	Met	Leu	Pro	
	65					70					75					8.0
45	Thr Leu	Leu	Ile	Leu	Glu	Trp	Val	Lys	Ala	Asp	Asn	Glu	Ser	Gly	Ser	
					85					90					95	
50	Leu	Cys	Leu	Thr	Lys	Glu	Asn	Leu	Thr	Asn	Phe	Gly	Leu	Asn	Thr	

Glu

100	105	110
700	702	770

_	Phe Leu	Ile	Ģlu	Ser	Leu	Gln	Asn	Ile	Ala	Gly	Ser	Glu	Cys	Leu	Asp
J	Беп		115	•				120					125		

- Ser Gln Arg Gln Glu Leu Thr Thr Arg Leu Asp Lys Ala Thr Met

 10 Ile

 130 135 140
- Leu Ser Leu Ser Val Pro Gln Ala Trp Leu Lys Tyr Gln Ala Thr
 15 Asn
 145 150 155
 160
- 20 Trp Thr Pro Pro Glu Phe Trp Asp Thr Gly Ile Thr Gly Phe Ile Leu 165 170 175
- 25 Asp Tyr Asn Val Tyr Ala Ser Gln Tyr Ala Pro His His Gly Asp Ser 180 185 190
- 30 Thr Gln Asn Val Ser Ser Tyr Gly Thr Leu Gly Phe Asn Leu Gly Ala
 195 200 205
- 35 Trp Arg Leu Arg Ser Asp Tyr Gln Tyr Asn Gln Asn Phe Ala Asp Gly 210 215 220
- 40 Arg Ser Val Asn Arg Asp Ser Glu Phe Ala Arg Thr Tyr Leu Phe Arg 225 230 235 240
- Pro Ile Pro Ser Trp Ser Ser Lys Phe Thr Met Gly Gln Tyr Asp Leu

 245
 250
 255

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	v	VO 200	5/0978	23									PCT	/ EP20 0	5/00397
								86/	370						
	Ser Glu	Ser	Asn	Leu	Tyr	Asp	Thr	Phe	His	Phe	Thr	Gly	Ala	Ser	Leu
				260					265					270	9
5			-: -:			_		_		_		~ 3	_		
	Ser Gln	Asp		Ser	Met	Leu	Pro		Asp	Leu	GIn	GTÀ	Tyr	Ala	Pro
			275					280					285		
10															
	Ile	Thr	Gly	Ile	Ala	Gln	Thr	Asn	Ala	Lys	Val	Thr	Val	Ala	Gln
		290					295					300			
1															
15	_	Arg	Val	Leu	Tyr	Gln	Thr	Thr	Val	Ala	Pro	Gly	Pro	Phe	Thr
	Ile 305					310					315				
	320					,					00	•			
20															
		Asp	Leu	Gly	Gln	Ser	Phe	Gln	Gly	Gln	Leu	Asp	Val	Thr	Val
	Glu				325					330	•				335
25															
	~ 7	~ 1	- 70	~]	-		~	rm 3	D.I	~ 3	** 7	61	2		~
	Glu Ile	Glu	Asp	_	Arg	Thr	Ser	Thr		GIn	Val	GTÀ	Ser		Ser
20				340			•		345					350	
30															
	Pro Gly	Tyr	Leu	Thr	Arg	Lys	Gly	Gln	Val	Arg	Tyr	ГÀг	Thr	Ser	Leu
	_		355					360					365		
35															
	_	Pro	Thr	Ser	Val	Gly	His	Asn	Asp	Ile	Asn	Asn	Pro	Phe	Phe
	Trp	370					375					380			
40															
					_		~ 7		_	_	-	7	_		
	Thr Gly	Ala	GLu	Ala	ser	Trp	GTA	Trp	ьeu	Asn	Asn	val	Ser	ьeu	туг
45	385 400					390					395				
. =															
	Gly	Gly	Met	Phe	Thr	Ala	Asp	Asp	Tyr	Gln	Ala	Ile	Thr	Thr	Gly

Ile

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	Gly Gly		Asn		Asn	Gln	Phe	Gly		Leu	Ser	Phe	Asp		Thr
5				420					425					430	
	Ala Tyr	Asp	Ala	Ser	Leu	Gln	Gln	Gln	Asn	Ser	Gly	Asn	Leu	Arg	Gly
10			435					440	,				445		
	Ser Gln	Tyr	Arg	Phe	Asn	Tyr	Ala	Lys	His	Phe	Glu	Ser	Thr	Gly	Ser
15		450					455					460			
	Ile Met	Thr	Phe	Ala	Gly	Tyr	Arg	Phe	Ser	Asp	Lys	Asp	Tyr	Val	Ser
20	465 480					470	*				475				
	0	G1		.	6	a	-	_	6 3		7				
25	Glu	GIU	Tyr	Leu	Ser 485	Ser.	Arg	Asn	GTA	490	GLu	Ser	Ile	Asp	Asn 495
					100					400					490
20	Lys Glu	Glu	Ser	Tyr	Val	Ile	Ser	Leu		Gln	Tyr	Phe	Glu		Leu
30				500					505					510	
· Y	Leu Ala	Asn	Ser	Tyr	Leu	Asn	Val	Thr	Arg	Asn	Thr	Tyr	Trp	Asp	Ser
35			515					520					525		
	Ser Gly	Asn	Thr	Asn	Tyr	Ser	Val	Ser	Val	Ser	Lys	Asn	Phe	Asp	Ile
40	2	530	i	•			535				٠	540			
		Phe	Lys	Gly	Ile	Ser	Ala	Ser	Leu	Ala	Val	Ser	Arg	Ile	Arg
45	Trp 545 560					550					555				
50	Asp Leu	Asp	Asp	Glu		Asn	Gln	Tyr	Tyr		Ser	Phe	·Ser ·	Leu	
					565					570					575

5	Gln Ser	Gln	Asn	Arg 580	Asn	Ile	Ser	Tyr	Ser 585	Met	Gln	Arg	Thr	Gly 590	Ser
10	Asn Asn	Thr	Ser 595	Gln	Met	Ile	Ser	Trp	Tyr	Asp	Ser	Ser	Asp 605	Arg	Asn
15	Ile Gly	Trp 610	Asn	Ile	Ser	Ala	Ser 615	Ala	Thr	Asp	Asp	Asn 620	Ile	Arg	Asp
20	Glu Arg 625 640	Pro	Thr	Leu	Arg	Gly 630	Ser	Tyr	Gln	His	Tyr 635	Ser	Pro	Trp	Gly
25	Leu Thr	Asn	Ile	Asn	Gly 645	Ser	Val	Gln	Pro	Asn 650	Gln	Tyr	Asn	Ser	Val 655
30	Ala Leu	Gly	Trp	Tyr 660	Gly	Ser	Leu	Thr	Ala 665	Thr	Arg	His	Gly	Val 670	Ala
35	His Asp	Asp	Tyr 675	Ser	Tyr	Gly	Asp	Asn 680	Ala	Arg	Met	Met	Val 685	Asp	Thr
40	Gly Gly	Ile 690	Ser	Gly	Ile	Glu	Ile 695	Asn	Ser	Asn	Arg	Thr 700	Val	Thr	Asn ·
45	Leu Met 705 720	Gly	Ile	Ala	Val	Ile 710	Pro	Ser	Leu	Ser	Asn 715	Tyr	Thr	Thr	Ser
50	Leu Asn	Arg	Val	Asn	Asn	Asn	Asp	Leu	Pro	Glu	Gly	Val	Asp	Val	Glu

Ser Val Ile Arg Thr Thr Leu Thr Gln Gly Ala Ile Gly Tyr Ala Lys Leu Asn Ala Thr Thr Gly Tyr Gln Ile Val Gly Val Ile Arg Gln Glu

Asn Gly Arg Phe Pro Pro Leu Gly Val Asn Val Thr Asp Lys Ala Thr 77 O

Gly Lys Asp Val Gly Leu Val Ala Glu Asp Gly Phe Val Tyr Leu 785 ·

Gly Ile Gln Glu Asn Ser Ile Leu His Leu Thr Trp Gly Asp Asn Thr

Cys Glu Val Thr Pro Pro Asn Gln Ser Asn Ile Ser Glu Ser Ala Ile

Ile Leu Pro Cys Lys Thr Val Lys

<210> 29 <211> 169 <212> PRT <213> Escherichia coli <400>

Leu Met Asn Thr Lys Gln Ser Val Ala Gln Leu Ala Val Pro His Arg

Lys Arg Leu Ser Ser Thr Met Val Val Ala Leu Leu Cys Val Val

	V	VO 200	05/0978	323				90/3	370				PCT	/ EP2 00	5/0039	72
		Gly	Ala	Val	Met	Ile	Asn			Asp	Phe	Pro	Ala	Thr	Ala	
	Ile		35					40					45			
5																
	Glu Ala		Asp	Pro	Gly	Ala	Ser	Ala	Phe	Pro	Thr	Phe	Tyr	Ala	Cys	
		50					55					60				
10	Leu	Ile	Val	Leu	Ala	Val	Leu	Leu	Val	Ile	Arq	Asp	Leu	Leu	Gln	٠
	Ala 65					70					75	-				80
15																
	Lys Lys	Pro	Ala	Ser	Cys	Ala	Asn	Ala	Gln	Glu	Lys	Pro	Ala	Phe	Arg	
	1				85					90	•		,		95	
20	Thr	Z\] =	Thr	Gly	Tla	73.7 =	7\] a	T hr	7\] =	Pho	Ψτ.75	Tlo	77a 7	7\ 7 -	M ~ 1-	
	Ser	ALG	1111	100	7.7.6	ALA	ALA	T111		FIIG	тут		vaı		Met	
25				100					105					110	•	
23		Cys	Gly	Tyr	Leu	Ile	Thr	Thr	Pro	Val	Phe	Leu	Ile	Val	Ile	
	Met		115					120				,	125			
30	en 1	_		~ 7	_					_		_			_	
	Tnr Leu		Met	Gly	Tyr	Arg		Trp	Val	Leu	Thr		GLy	Ile	Ala	
		130					135					140				
35		Leu	Thr	Ala	Ile	Leu	Trp	Leu	Leu	Phe	Val	Glu	Ala	Leu	Gln	
	Val 145					150					155					
40	160															
	Pro	Leu	Pro	Val	Gly 165	Thr	Phe	Phe	Glu							
45	.0.0	\.			0.1		•									

	Ala Ala	Ser	Tyr	Pro	Thr	Lys	Gln	Ile	Glu	Leu	Val	Val	Pro	Tyr	Ala	
5	AIA			20					25					30		
	Gly Lys	Gly	Gly	Thr	Asp	Leu	Vаl	Ala	Arg	Ala	Phe	Ala	Asp	Ala	Ala	
10			35				ų.	40					45			
	Asn Gly	His	Leu	Pro	Val	Ser	Ile	Gly	Val	Ile	Asn	Lys	Pro	Gly	Gly	
15		50					55					60				
	Gly Tyr	Ala	Ile	Gly	Leu	Ser	Glu	Ile	Ala	Ala	Ala	Arg	Pro	Asn	Gly	
20	65					70					75				,	80
	Lys Gly	Ile	Gly	Leu	Gly	Thr	Val	Glu	Leu	Thr	Thr	Leu	Pro	Ser	Leu	
25					85					90					95	
	Met Asn	Val	Arg	Phe	Lys	Thr	Ser	Asp	Phe	Lys	Pro	Ile	Ala	Arg	Leu	
30	23.011			.100					105					110		
		Asp	Pro	Ala	Ala	Ile	Thr	Val	Arg	Ala	Asp	Ala	Pro	Trp	Asn	
35	Ser		115					120	,				125			
		Glu	Glu	Phe	Met	Ala	Tyr	Ser	Lys	Ala	Asn	Pro	Gly	Ļys	Val	
40	Ar.g	130					135					140				
		Gly	Asn	Ser	Gly	Thr	Gly	Ala	Ile	rrp	His	Leu	Ala	Ala	Ala	
45	Ala 145 160					150					155					

Leu Glu Asp Lys Thr Gly Thr Lys Phe Ser His Val Pro Tyr Asp

165

170

175

50

Gly

	Ala Val	Ala	Pro	Ala	Tle	Thr	Gly	Leu	Leu	Gly	Gly	His	Ile	Glu	Ala
5	Vul			180					185					190	
	Ser	Val	Ser	Pro	Gly	Glu	Val	Ile	Asn	His	Val	Asn	Gly	Gly	Lys
10	Leu		195					200					205		
	т	mb	Tou	V2]	Val	Mot	Δla	\(\frac{1}{2}\) sn	Glu	Ara	Met	Tivs	Thr	Met	Pro
	Lys		ьeu	Val	Val	Met.		дър	GIG	1119	1100				· .
15		210					215					220			
		Pro	Thr	Leu	Lys	Glu	Lys	Gly	Val	Asp	Leu	Ser	Ile	Gly	Thr
20	Trp 225 240					230					235				
	240														
0.5		Gly	Leu	Ile	·Val	Ser	Gln	Lys	Thr	Pro	Gln	Asp	Val	Val	Asp
25	Val				245					250					255
	Leu	Ala	Lys	Ala	Ala	· Lys	Glu	Thr	Ala	Glu	Glu	Pro	Ala	Phe	Gln
30	Asp		_	260					265					270	
														w 7	77.7
35	Ala Ser		Gln	Lys	Leu	Asn	Leu	Asn	Tyr	Ala	Trp	Leu			Ala
			275	•				280					285		
	Phe	Gln	Thr	Gln	Ile	ser	Glu	Gln	Glu	. Lys	Tyr	Phe	: Asp	Glu	Leu
40	Leu	290)				295					300)		
15	Thr		g Leu	ı Gly	. Leu	Lys 310		;							

<210> 31 <211> 722 <212> PRT <213> Escherichia coli <400> 31

310

50

45

	93/370
	73/3/1

		Leu	Arg	Trp	Lys	Arg	Cys	Ile	Ile	Leu	Thr	Phe	Ile	Ser	Gly	
-	Ala 1				5					10		,			15	
5	Ala Pro	Phe	Ala	Ala	Pro	Glu	Ile	Asn		Lys	Gln	Asn	Glu		Leu	
				20					25					30		
10	Asp Gly	Leu	Gly	Ser	Gln	Ala	Ala	Gln	Gln	Asp	Glu	Gln	Thr	Asn	Lys	
	Ç		35		4			40					45			
15	-	Ser	Leu	Lys	Glu	Arg	Gly	Ala	Asp	Tyr	Val	Ile	Asn	Ser	Ala	
	Thr	50					55					60				
20		Gly	Phe	Glu	Asn	Leu	Thr	Pro	Glu	Ala	Leu	Glu	Ser	Gln	Ala	
	Arg 65					70					75~		-			. 80 ·
25		Tyr	Leu	Gln	Ser	Gln	Ile	Thr	Ser	Thr	Ala	Gln	Ser	Tyr	Ile	
	Glu				85					90					95	
30	-	Thr	Leu	Ser	Pro	Tyr	Gly	Lys	Val	Arg	Leu	As n	Leu	Ser	Ile	
	Gly			100					105					110		
35	Gln	Gly	Gly	Asp	Leu	Asp	Gly	Ser	Ser	Ile	Asp	Tyr,	Phe	Val	Pro	
	Trp		115					120					125			
40	Tyr	Asp	Asn	Gln	Thr	Thr	Val	Tyr	Phe	Ser	Gln	Phe	Ser	Ala	Gln	
	Arg	130					135					140				
45	Lvs	Glu	Asp	Arq	Thr	Ile	Gly	Asn	Ile	Gly	Leu	Glу	Val	Arg	Tyr	
	Asn 145		r	,		150	-			-	155	_				
50	160											*				

,	· · · · ·										· -,	_ 0 0 0, 0 .	· · · -	
					9	4/370								
												•		
Phe As	p Lys	Tyr	Leu	Leu	Gly	Gly	Asn	Ile	Phe	Tyr	Asp	Tyr	Asp	

Phe 165 170 175

Thr Arg Gly His Arg Arg Leu Gly Leu Gly Ala Glu Ala Trp Thr
Asp
180
185
190

Tyr Leu Lys Phe Ser Gly Asn Tyr Tyr His Pro Leu Ser Asp Trp Lys

195
200
205

Asp Ser Glu Asp Phe Asp Phe Tyr Glu Glu Arg Pro Ala Arg Gly
Trp
210
215
220

Asp Ile Arg Ala Glu Val Trp Leu Pro Ser Tyr Pro Gln Leu Gly Gly 225 230 235

25

30

35

40

45

50

Lys Ile Val Phe Glu Gln Tyr Tyr Gly Asp Glu Val Ala Leu Phe Gly

245
250
255

Thr Asp Asn Leu Glu Lys Asp Pro Tyr Ala Val Thr Leu Gly Leu Asn 260 265 270

Tyr Gln Pro Val Pro Leu Leu Thr Val Gly Thr Asp Tyr Lys Ala Gly 275 280 280 285

Thr Gly Asp Asn Ser Asp Val Ser Ile Asn Ala Thr Leu Asn Tyr Gln 290 295 300

Phe Gly Val Pro Leu Lys Asp Gln Leu Asp Ser Asp Lys Val Lys Ala 305 310 315

			02,0270	, _				95/	370					1,2120	
	Ala Asn	His	Ser	Leu	Met	Gly	Ser	Arg	Leu	Asp	Phe.	Val	Glu	Arg	Asn
5					325					330					335
	Phe Leu	Ile	Val	Leu	Glu	Tyr	Lys	Glu	Ļуs	Asp	Pro	Leu	Asp	Val	Thr
.10				340					345					350	
	Trp Lys	Leu	Lys	Ala	Asp	Ala	Thr	Asn	Glu	His	Pro	Glu	Cys	Val	Ile
15	.4		355					360				•	365		
	Asp Ile	Thr	Pro	Glu	Ala	Ala	Val	Gly	Leu	Glu	Lys	Cys	Lys	Trp	Thr
20	110	370					375					380			
	Asn Gln	Ala	Leu	Ile	Asn	His	His	Tyr	Lys	Ile	Val	Ala	Ala	Ser	Trp
25	385 400					390					395				
	Ala Glu	Ĺys	Asn	Asn	Ala	Ala	Arg	Thr	Leu	Val	Met	Pro	Val	Ile	Lys
30					405					410	-				415
	Asn Pro	Thr	Leu	Thr	Glu	Gly	Asn	Asn	Asn	His	Trp	Asn	Leu	Val	Leu
35				420		•			425					430	
	Ala Thr	Trp	Gln	Tyr	Ser	Ser	Asp	Gln	Ala	Glu	Gln	Glu	Lys	Leu	Asn
40			435					440					445		
	Trp Asn	Arg	Val	Arg	Leu	Ala	Leu	Glu	Asp	Glu	Lys	Gly	Asn	Arg	Gln

Ser Gly Val Val Glu Ile Thr Val Gln Gln Asp Arg Lys Ile Glu Leu 470 475

460

455

45

	Ile Glu	Val	Asn	Asn	Ile	Ala	Asn	Pro	Glu	Glu	Asn	Asn	His	Ser	His
5					485					490					495
	Ala	Ser	Ala	Gln	Ala	Asp	Gly	Val	Asp	Gly	Val	Va·l	Met	Asp	Leu
10	Asp			500					505					510	
	Val	Thr	Asp	Ser	Phe	Gly	Asp	Asn	Thr	Asp	Arg	Asn	Gly	Asp	Ala
15	Leu		515					520					525	_	
	Pro	Glu	Asp	Asn	Leu	Thr	Pro	Gln	Leu	Tyr	Asp	Ala	Gln	Asp	Lvs
20	Arg	530					535					540		-	2
	Val	Thr	Leu	Thr	Asn	Lys	Pro	Cys	Ser	Thr	Asp	Asn	Pro	Cys	Val
25	Phe 545 560					550					555			-	
	Ile	Ala	Lys	Gln	Asp	Lys	Glu	Lys	Gly	Thr	Val	Thr	Leu	Ser	Ser
30	Thr				565					570					575
	Leu	Pro	Gly	Thr	Tyr	Arg	Trp	Lys	Ala	Lys	Ala	Ala	Pro	Tyr	Asp
35	Asp			580					585					590	-
	Ser	Asn	Tyr	Val	Asp	Val	Thr	Phe	Leu	Gly	Ala	Glu	Ile	Glv	Gl v
40	Leu		595					600		-			605	0 -1	011
	Asn	Ala	Phe	Tle	Тυν	Δκα	V = 1	Cl u	7 7 ~	71 -	Tvic	Dage	C	77	-
45	Ile	610			- y -	231 Y	615	ЭтХ	nia	лта	пÀЗ	620	ser	ASN	ьеи
							- — -					220			
50	Gly Phe	Lys	Asp	Lys	Glu	Pro	Leu	Рŗо	Ser	Thr	Thr	Phe	Ile	Asp	Leu

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97/370 Tyr Gly Ala Thr Thr Ile Lys Thr Val Ser Ser Arg Ser Lys Asn Leu Thr Lys Arg Trp Cys Ser Thr Thr Thr Ser Gly Asn Leu Pro Ala Arg Ala Ser Met Val Ser Gly Cys Thr Gly Glu His Ser Asn Glu Asp. Ile Val Ile Pro Ala Thr Asn Arg Glu Ala Ala Gln Thr Tyr Gly Ala 690 Gln Ala Gly Asp Gly Leu Gln Gly Tyr Gly Leu Arg Val Leu Tyr Lys Lys <210> 32 <211> 319 <212> PRT <213> Escherichia coli <400> Met Lys Gln Asp Lys Arg Arg Gly Leu Thr Arg Ile Ala Leu Ala Leu Ala Leu Ala Gly Tyr Cys Val Ala Pro Val Ala Leu Ala Glu Asp Ser

Ala Trp Val Asp Ser Gly Glu Thr Asn Ile Phe Gln Gly Thr Ile

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Trp Leu Tyr Ser Glu Gly Gly Ser Ala Thr Thr Asp Ala Asp Arg Val Thr Leu Thr Ser Asp Leu Lys Gly Ala Arg Pro Gln Gly Met Lys Arq Thr Ser Val Phe Thr Arg Val Ile Asn Ile Gly Asp Thr Glu Gly Asp Val Asp Leu Gly Gly Leu Gly Asp Asn Ala Lys Thr Ile Asp Thr Ile Arg Trp Met Ser Tyr Lys Asp Ala Gln Gly Gly Asp Pro Lys Glu Leu Ala Thr Lys Val Thr Ser Tyr Thr Leu Thr Asp Ala Asp Arg Gly Arg 1.35 Tyr Ile Gly Ile Glu Ile Thr Pro Thr Thr Gln Thr Gly Thr Pro Asn Val Gly Thr Ala Leu His Leu Tyr Asp Val Ser Thr Ala Ser Gly Ġly Gly Ser Asp Ser Asp Asn Val Ala Pro Gly Pro Val Val Asn Gln Asn Leu Lys Val Ala Ile Phe Val Asp Gly Thr Ser Ile Asn Leu Ile Asn

		Ser	Thr	Pro	Ile	Glu	Leu	Gly	Lys	Thr	Tyx	Val	Ala	Lys	Leu
	Tyr	210					215					220			
5												-		_	
	Ser Asp	Asp	Glu	Asn	Lys	Asn	Gly	Lys	Phe	Asp	Ala	Gly	Thr	Asp	Ala
10	225 240					230					235				
10	240														
		Thr	Ala	Asn	Tyr	Asp	Phe	Arg	Trp	Val	Leu	Ser	Gly	Ser	Ser .
15	Gln				245					250					255
	Gln Asn	Leu	Gly	Thr	Ser	Gly	Gly	Ile	Val	Asn	Ser	Ser	Phe	Asp	Asn
20	11011			260		٠			265			,		270	
	_	-	r. 7	~ 1 -	Dana	71 7 -	mb ~	7 an	λαn	Glu	7A 71 ==	Ara	Thr	Asn	T.e.11
	Asn Asn	Leu		TTE	PIO	Ата	TIIT		Asp	OLU	Ala	117.3		11011	200
25			275					280					285		
	Gly	Pro	Ala	Arg	Asp	Gly	Lys	Glu	Ala	Leu	Ser	Ile	Pro	Thr	Asn
30	Gly	290					295					300			
		Gly	Val	Gln	Gly	Tyr 310	Lys	Leu	His	Ile	Ile 315	Tyr	Lys	His	Lys
35	305					210					313				
	<21	0>	33 <	211>	62	9 <2	12>	PRT	<21	3>	Esche	∍ric	hia	coli	<400>
	33									•					
40	Met His		Lys	Val	Leu	Thr	Leu	Ser	Leu	Leu	Ala	Leu	. Cys	Val	Ser
	1				5					10					15
4.5		~ 7	** - 7	7. 1 _	71 7	71		, mb r	Pho	Ner	. Aen	Zlan	. Asn	Tle	Ala
45	Ser Leu		vaı		Ald	I ASI.	г тат			MSI.	n Asn	2101	1101.		
				20					25					30	
50	Ser	· Phe	Asp	asp	Thr	Asr	n Ser	: Thr	: Ile	. Val	Leu	Lys	: Asp	Arg	Arg
	Thr		- L	_											

Thr

35 40 45

Asn His Pro Ile Thr Pro Gln Glu Leu Phe Phe Leu Thr Leu Pro 5 Asp 50 55 60

Glu Thr Lys Ile His Thr Ala Asp Phe Lys Ile Lys His Ile Lys
10 Lys
65 70 75 80

Gln Asp Asn Ala Ile Val Ile Asp Phe Thr Arg Pro Asp Phe Asn
Val
85 90 95

Thr Val Gln Leu Asn Leu Val Lys Gly Lys Tyr Ala Ser Ile Asp 20 Tyr 100 105 110

Thr Ile Ala Ala Val Gly Gln Pro Arg Asp Val Ala Lys Ile Thr 25 Phe 115 120 125

Phe Pro Thr Lys Lys Gln Phe Gln Ala Pro Tyr Val Asp Gly Ala 30 Ile 130 135 140

Thr Ser Ser Pro Ile Ile Ala Asp Ser Phe Phe Ile Leu Pro Asn
Lys
145
160

40 Pro Ile Val Asn Thr Tyr Ala Tyr Glu Ala Thr Thr Asn Leu Asn Val
. 165 170 175

45 Glu Leu Lys Thr Pro Ile Gln Pro Glu Thr Pro Val Ser Phe Thr
Thr
180 185 190

50 Trp Phe Gly Thr Phe Pro Glu Thr Ser Gln Leu Arg Arg Ser Val Asn

195	200	205

5	Gln His	Phe 210	Ile	Asn	Ala	Val	Arg 215	Pro	Arg	Pro	Tyr	Lys 220	Pro	Tyr	Leu
10	Tyr Gln 225 240	Asn	Ser	Trp	Met	Asp 230	Ile	Gly	Phe	Phe	Thr 235	Pro	Tyr	Thr	Glu
15	Asp Gly	.Val	Leu	Gly	Arg 245	Met	Asp	Glu	Trp	Asn 250	Lys	Glu	Phe	Ile	Ser 255
20	Arg Asp	Gly	Val	Ala 260	Leu	Asp	Ala	Phe	Leu 265	Leu	Asp	Asp	Gly	Trp 270	Asp
25	Leu Ser	Thr	Gly 275	Arg	Trp	Leu	Phe	Gly 280	Pro	Ala	Phe	Ser	Asn 285	Gly	Phe
30	Lys Trp	Val 290.	Arg	Glu	Lys	Ala	Asp 295		Leu	His	Ser	Ser 300	Val	Gly	Leu
35	Leu Phe 305 320	Ser	Pro	Trp	Gly	Gly 310	Tyr	Asn	Lys	Pro	Gln 315	Arg	Arg	Ser	Arg
40	Ala Ala	Cys	Lys	Arg	Val	Trp	Val	Arg	Asn	Arg 330	Gly	Arg	Gln	Ala	Gly 335
45	Phe Leu	Gly	Ser	Glu 340	Leu	Leu	Lys	Asn	Phe	Asn	Glu	Gln	Ile	Ile 350	Asn

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	V	V O 200	5/09782	23				PCT/EP2005/00393							
	Ile	Lys	Asn	Glu	His	Ile	Thr	Ser	Phe	Lys	Leu	Asp	Gly	Met	Gly
	Asn		355					360					365		
5	Ala Ala	Ser	Ser	His	Ile	Lys	Gly	Ser	Pro	.Phe	·Ala	Ser	Asp	Phe	Asp
	ALA	370					375					380			
10	Ser Phe	Ile	Ala	Leu	Leu	His	Asn	Met	Arg	Arg	Ala	Asn	Pro	Asn	Leu
1.7	385 400					390					395		ı		
15								_	~ 3	_	_	~		-	D.1
	Ile Tyr	Asn	Leu	Thr	Thr	Gly	Thr	Asn	Ala		Pro	Ser	Trp	Leu	•
20					405				_	410		•			415
	Ala Pro	Asp	Ser	Ile	Trp	Arg	Gln	Gly	Asp	Asp	Ile	Asn	Leu	Tyr	Gly
25	11.0			420	•				425					430	
	Gly Tyr	Thr	Pro	Val	Gln	Gln	Trp	Ile	Thr	Tyr	Arg	Asp	Ala	Glu	Thr
30	тут		435					440					445		
	Arg Met	Ser	Ile	Val	Arg	Lys	Gly	Pro	Leu	Phe	Pro	Leu	Asn	Ser	Leu
35	nec	450					455					460			
	_	His	Gly	Ile	Val	Ser	Ala	Glu	Asn	Ala	Tyr	Tyr	Gly	Leu	Glu
40	Lys 465 480					470					475				
4.5	Val Ala	Gln	Thr	Asp		Asp	Phe	Ala	Asp		Val	Trp	Ser	Tyr	
45					485					490					495

Thr Gly Thr Gln Leu Gln Glu Leu Tyr Ile Thr Pro Ser Met Leu Asn

		Val	Lys	Trp	Asp	Thr	Leu	Ala	Lys	Ala	Ala	Lys	Trp	Ser	Lys
5	Glu		515					520					525		
J	Asn	Ala	Ser	Val	Leu	Val	Asp	Thr	His	Trp	Tle	Glv	Glv	Asp	Pro
	Thr	530	201	• • •	200	,	535					540		1101	
10							·								
	Ala Ile	Leu	Ala	Val	Tyr		Trp	Ala	Ser	Trp	Ser	Lys	Asp	Lys	Ala
15	545 560					550					555				
	Tou	Clv	Leu	7\ r. c.	7) en	Dro	Ser	7\ e.p.	Twa	Pro	Cln	Thγ	Ф.,,,	Πτ.π	Tou
20	Asp	GIY	пец	ALG	565	FIO	Ser	vsb	пур	570	Gili	7117	тут	тут	575
20					000					3,0		-			070
	Leu Ser	Ala	Lys	Asp	Phe	Glu	Ile	Pro ·	Ala	Gly	Asn	Ala	Ala	Gln	Phe
25				580			*		585					590	
		Lys	Ala	Val	Tyr	Gly	Ser	Asn	Lys	Thr	Val	Pro	Val	Glu	Tyr
30	Lys		595					600					605		
	Asn	Ala	Thr	Val	Ile	Thr	Leu	Gln	Pro	'Leu	Glu	Thr	Leu	Val	Phe
35	Glu	610					615					620	-		
							•								
	Ala 625	Val	Thr	Ile	Asn										
40	-01		24 - 6	711	1 77 7	70 - (2105	DDr	n /01	1.25	T l		-1-2-	- 7	
	<400		34 <2 34	7.T.T.>	17.	18 <2	21 <i>2></i>	PR.	Ľ <z.< td=""><td>13></td><td>ESCI</td><td>nerio</td><td>cnia</td><td>COT</td><td>L</td></z.<>	13>	ESCI	nerio	cnia	COT	L
45	Met Tyr	Asn	Lys	Ile	Phe	Lys	Val	Ile	Trp	Asn	Pro	Ala	Thr	Gly	Ser
	1				5					10		٠	-		15
50	Thr	Val	Ala	Ser	Gļu	Thr	Ala	Lys	Ser	Arg	Gly	Lys	Lys	Ser	Gly

Arg

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20	25	3
		_

0

Ser Lys Leu Leu Ile Ser Ala Leu Val Ala Gly Gly Leu Leu Ser 5 Ser 40 45

Phe Gly Ala Ser Ala Asp Asn Tyr Thr Gly Gln Pro Thr Asp Tyr 10 Gly $50 \hspace{1.5cm} 55 \hspace{1.5cm} 60$

Asp Gly Ser Ala Gly Asp Gly Trp Val Ala Ile Gly Lys Gly Ala

15 Lys
65 70 75 80

Ala Asn Thr Phe Met Asn Thr Ser Gly Ala Ser Thr Ala Leu Gly
Tyr
85
90
95

Asp Ala Ile Ala Glu Gly Glu Tyr Ser Ser Ala Ile Gly Ser Lys

Thr

100 105 110

Leu Ala Thr Gly Gly Ala Ser Met Ala Phe Gly Val Ser Ala Lys 30 Ala 115 120 125

Met Gly Asp Arg Ser Val Ala Leu Gly Ala Ser Ser Val Ala Asn
35 Gly
130 135 140

Asp Arg Ser Met Ala Phe Gly Arg Tyr Ala Lys Thr Asn Gly Phe
40 Thr
145 150 155

50 Ala Leu Gly Asn Thr Ala Lys Ala Tyr Glu Ile Met Ser Ile Ala Leu

190

103/3/10

180

Gly Asp Asn Ala Asn Ala Ser Lys Glu Tyr Ala Met Ala Leu Gly 5 Ala

195 200 205

Ser Ser Lys Ala Gly Gly Ala Asp Ser Leu Ala Phe Gly Arg Lys

10 Ser
210 215 220

Thr Ala Asn Ser Thr Gly Ser Leu Ala Ile Gly Ala Asp Ser Ser
Ser
225
240
235

20 Ser Asn Asp Asn Ala Ile Ala Ile Gly Asn Lys Thr Gln Ala Leu Gly
245 250 255

25 Val Asn Ser Met Ala Leu Gly Asn Ala Ser Gln Ala Ser Gly Glu Ser
260 265 270

30 Ser Ile Ala Leu Gly Asn Thr Ser Glu Ala Ser Glu Gln Asn Ala Ile 275 280 285

35 Ala Leu Gly Gln Gly Ser Ile Ala Ser Lys Val Asn Ser Ile Ala Leu 290 295 300

Gly Ser Asn Ser Leu Ser Ser Gly Glu Asn Ala Ile Ala Leu Gly Glu 305 310 315

Gly Ser Ala Ala Gly Gly Ser Asn Ser Leu Ala Phe Gly Ser Gln Ser
325
330
335

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345 345 346 347 348		Arg Ala	Ala	Asn	Gly	Asn	Asp	Ser	Val	Ala	Ile	Gly	Val	Gly	Ala	Ala
Ala Thr Asp Asn Ser Val Ala Ile Gly Ala Gly Ser Thr Thr Asp Ala Ala Gly Ser Thr Thr Asp Ala Ala Gly Ash Ash Ash Ash Ser Ash Thr Val Ser Val Gly Ash Ser Ala Thr Lys Arg Lys Ile 370				-	340					345					350 ·	
10	5		Thr	Asp	Asn	Ser	Val	Ala	Ile	Gly	Ala	Gly	Ser	Thr	Thr	Asp
Ser Asn Thr Val Ser Val Gly Asn Ser Ala Thr Lys Arg Lys Ile Asn		ALA		355					360					365	•	
370 375 380 15	10	Se <i>x</i>	Asn	Thr	Val	Ser	Val	Gly	Asn	Şer	Ala	Thr	Lys	Arg	Lys	Ile
Asn Met Ala Ala Gly Ala Ile Ser Asn Thr Ser Thr Asp Ala Ile Asn 385 400 20 20		Val	370					375					380			
Assn 38 5 4000 20 Ser Gln Leu Tyr Thr Ile Ser Asp Ser Val Ala Lys Arg Leu	15	Asn	Met	Ala	Δla	Glv	Δla	Tle	Ser	Asn	Thr	Ser	Thr	Asn	70.1 a	Tle
20		Asn 385	1100	1114	7114	С±у		110	DCI	71511	1111		X 11.L	дзр	Ата	TTC
25	20	400														
25				Gln	Leu	Tyr	Thr	Ile	Ser	Asp	Ser	Val	Ala	Lys	Arg	Leu
130	25					405					410					415
30			Gly	Ala	Thr	Val	Gly	Ser	Asp	Gly	Thr	Val	Thr	Ala	Val	Ser
35 435 440 445 445 445 445 445 445 455 470 475 475 475 460 475 475 475 475 475 475 475 475 475 475	30	± y ±			420					425		•			430	
35 445 4			Leu	Arg	Ser	Gly	Thr	Tyr	Asn	Asn	Val	Gly	Asp	Ala	Leu	Ser
Ile Asp Asn Asn Thr Leu Gln Trp Asn Lys Thr Ala Gly Ala Phe Ser 450	35	GTĀ		435					440	•	٠			445		
450 455 460 460 460 460 460 460 460 460 460 460		Ile	Asp	Asn	Asn	Thr	Leu	Gln	Trp	Asn	Lys	Thr	Ala	Gly	Ala	Phe
Ala Asn His Gly Ala Asn Ala Thr Asn Lys Ile Thr Asn Val Ala Lys 465 470 475	4.0	Ser	450					455					460			
Lys 465 470 475	40	73.1 - 2	7) en	Пie	G1 w	70.7 =	Z) s n	7.1.5	Thr	7) = n	T.ve	Tla	Thr	7) e n	۲ <i>۲</i> م ۲	7.1.5
	45	Lys 465		1113	GIY	AIG		ALA	1111	ASII	, nys		T 111	ASII	vai	AIA
Gly Thr Val Ser Ala Thr Ser Thr Asp Val Val Asn Gly Ser Gln		Gly	Thr	Val	Ser	Ala	Thr	Ser	Thr	Asp	Val	Val.	Asn	Gly	Ser	Gln
Leu 50 485 490 495	50									<u>.</u>				-		

								107/3	70						
	Tyr Ser	Asp	Leu	Gln	Gln	Asp	Ala	Leu	Leu	Trp	Asn	Gly	Thr	Ala	Phe
5				500	7				505					510	
	Ala Ala	Ala	His	Gly	Thr	Glu	Ala	Thr	Ser	Lys	Ile	Thr	Asn	Val	Thr
10			515					520					525		
	Gly Leu	Asn	Leu	Thr	Ala	Gly	Ser	Thr	Asp	Ala	Val	Asn	Gly	Ser	Gln
15		530					535					540			
	Lys Thr	Thr	Thr	Asn	Asp	Asn	Val	Thr	Thr	Asn	Thr	Thr	Asn	Ile	Ala
20	545 560					550					555				
	Asn Gly	Thr	Thr	Asn	Ile	Thr	Asn	Leu	Thr	Asp	Ala	Val	Asn	Gly	Leu
25					565			,		570					575
	Asp Ala	Asp	Ser	Leu	Leu	Trp	Asn	Lys	Ala	Ala	Gly	Ala	Phe	Ser	Ala
30				580					585					590	
	His Asn	Gly	Thr	Glu	Ala	Thr	Ser	Lys	Ile	Thr	Asn	Val	Thr	Ala	Gly
35			595					600					605		
	Leu Thr	Thr	Ala	Gly	Ser	Thr	Asp	Ala	Val	Asn	Gly	Ser	Gln	Leu	Lys
40		610		-			615					620			
	Thr	Asn	Asp	Asn	Val	Thr	Thr	Asn	Thr	Thr	Asn	Ile	Ala	Thr	Asn

Thr Asn Ile Thr Asn Leu Thr Asp Ala Val Asn Gly Leu Gly Asp

50 Asp

645 650 655

5	Ser Gly	Leu	Leu	Trp	Asn	Lys	Thr	Ala	Gly	Ala	Phe	Ser	Ala	Ala	His
	-			660					665					670	
10 15	Thr Thr	Asp	Ala	Thr	Ser	Lys	Ile	Thr	Asn	Val	Thr	Ala	Gly	Asn	Leu
	T11T		675					680					685		
	Ala	Gly	Ser	Thr	Asp	Ala	Val	Asn	Gly	Ser	Gln	Leu	Lys	Thr	Thr
	Asn	690					695					700			
	Asp	Asn	Val	Thr	Thr	Asn	Thr	.Thr	Asn	Tle	Ala	Thr	Asn	Thr	Thr
20	Asn 705	11011				710			11011		715	2112			1111
	720														
25		Thr	Asn	Leu	Thr	Asp	Ala	Val	Asn	Gly	Leu	Gly	Asp	Asp	Ser
	Leu	_			725					730	-				735
30	Leu	Trp	Asn	Lys	Thr	Ala	Gly	Ala	Phe	Ser	Ala	Ala	His	Gly	Thr
	Asp			740					745					750	
35	Ala	Thr	Ser	Lvs	Ile	Thr	Asn	Val	Ĺvs	Ala	Glv	Asp	Leu	Thr	Ala
	Gly		755	- <i>x</i> ~				760	-1-	4 4 4 4 4	<i>1</i>	1101	765		73.E.Q
		_,				_					<u>.</u>				
40	Ser Asn	770	Asp	Ala	Val	Asn	Gly 775	Ser	Gln	Leu	Lys		Thr	Asn	Asp
		770					775					780			
45	Val Asn 785 800	Ser	Thr	Asn	Thr	Thr	Asn	Ile	Thr	Asn	Leu	Thr	Asp	Ala	Val
						790					795				
50	Glv	T,e11	Glu	Aan	Asn	Ser	T.e.;	Leu	甲ェハ	Δen	T.ve	Thr	7 <u>)</u>] =	G1 11	70.7 -2
50	Phe	⊥ieu	Ψ¥	1.70 D	133 D	DOT	ъси	ചായ	115	UOII	пλο	TIIT	∆T.a	атЛ	пта

805 810 815

Ser Ala Ala His Gly Thr Asp Ala Thr Ser Lys Ile Thr Asn Val 5 Lys 820 825 830

Ala Gly Asp Leu Thr Ala Gly Ser Thr Asp Ala Val Asn Gly Ser 10 Gln 835 840 845

Leu Lys Thr Thr Asn Asp Asn Val Ser Thr Asn Thr Thr Asn Ile

15 Thr

850 855 860

Asn Leu Thr Asp Ser Val Gly Asp Leu Lys Asp Asp Ser Leu Leu 20 Trp 865 870 875 880

25 Asn Lys Ala Ala Gly Ala Phe Ser Ala Ala His Gly Thr Glu Ala Thr 885 890 895

30 Ser Lys Ile Thr Asn Leu Leu Ala Gly Lys Ile Ser Ser Asn Ser Thr 900 905 910

35 Asp Ala Ile Asn Gly Ser Gln Leu Tyr Gly Val Ala Asp Ser Phe Thr
915 920 925

40 Ser Tyr Leu Gly Gly Gly Ala Asp Ile Ser Asp Thr Gly Val Leu Ser 930 935 940

45 Gly Pro Thr Tyr Thr Ile Gly Gly Thr Asp Tyr Thr Asn Val Gly Asp 945 950 955

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	(WO 2 00:		110/370						PCT/EP2005/003972			3972		
								110,0,	o .						
	Ala Ala	Leu .	Ala A	Ala :	Ile 1	Asn :	Thr Se	er Ph	ne Se	er Th	nr Se	er Leu	ı Gly	/ Asp	
	1114			. 9	965				97	70				975	5
5	Leu Ile	Leu			Ala '	Thr A	Ala Gl			ne Se	er Al	La Lys		_	7
				980				- 98	35			•	990)	
10	Asn Ser	Asn .	Ala I	Pro S	Ser '	Val 1	[le Th	nr A	/sp /	/al <i>I</i>	Ala <i>A</i>	Asn Gl	Ly <i>P</i>	Ala V	7al
			995				10	000				1(05		
15	Ser	Thr 1010		Ser	Asp	Ala	Ile 1015	Asn	Gly	Ser	Gln	Leu 1020	Tyr	Gly	Val
20	Ser	Asp 1025	_	Ile	Ala	Asp	Ala 1030	Leu	Gly	Gly	Asn	Ala 1035	Val	Val	Asn
25	Thr	Asp 1040	_	Ser	Ile	Thr	Thr 1045	Pro	Thr	Tyr	Ala	Ile 1050	Ala	Gly	Gly
30	Ser	Tyr 1055		Asn	Val	Gly	Asp 1060	Ala	Leu	Glu	Ala	Ile 1065	Asp	Thr	Thr
•	Leu	Asp 1070	_	Ala	Leu	Leu	Trp 1075	Asp	Thr	Thr	Ala	Asn 1080	Gly	Gly	Asn
35	Gly	Ala 1085	Phe	Ser	Ala	Ala	His 1090	Gly	Lys	Asp	Lys	Thr 1095	Ala	Ser	Val
40	Ile	Thr 1100	Asn	Val	Ala	Asn	Gly 1105	Ala	Val	Ser	Ala	Thr 1110	Ser	Asn	Asp
45	Ala	Ile 1115	Asn	Gly	Ser	Gln	Lец 1120	Tyr	Ser	Thr	Asn	Lys 1125	Туr	Ile	Ala

Asp Ala Leu Gly Gly Asp Ala Glu Val Asn Ala Asp Gly Thr Ile

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	1 1 1/-3 / 11

	Thr	Ala 1145	Pro	Thr	Tyr	Thr	Ile 1150	Ala	Asn	Thr	Asp	Tyr 1155	Asn	Asn	Val
5	Gly	Glu 1160		Leu	Asp ·	Ala	Leu 1165	Asp	Asn	Asn	Ala	Leu 1170	Leu	Trp	Asp
10	Glu	Asp 1175	Ala	Gly	Ala	Tyr	Asn 1180	Ala	Ser	His	Asp	Gly 1185	Asn	Ala	Ser
15	Lys	Ile 1190	Thr	Asn	Val	Ala	Ala 1195	Gly	Asp	Leu	Ser	Thr 1200	Thr	Ser	Thr
	Asp	Ala 1205	Val	Asn	Gly	Ser	Gln 1210	Leu	Asn	Ala	Thr	Asn 1215	Ile	Leu	Val
20	Thr	Gln 1220	Asn	Ser	Gln	Met	Ile 1225	Asn	Gln	Leu	Ala	Gly 1230	Asn	Thr	Ser
25	Glu	Thr 1235	Tyr	Ile	Glu	Glu	Asn 1240	Gly	Ala	Gly	Ile	Asn 1245	Tyr	Val	Arg
30	Thr	Asn 1250	Asp	Ser	Gly	Leu	Ala 1255	Phe	Asn	Asp	Ala	Ser 1260	Ala	Ser	Gly
35	Ile	Gly 1265	Ala	Thr	Ala	Val	Gly 1270	Tyr	Asn	Ala	Val	Ala 1275	Ser	His	Ala
	Ser	Ser 1280	Val	Ala	Ile	Gly	Gln 1285	Asp	Ser	Ile	Ser	Glu 1290		Asp	Thr
40	Gly	Ile 1295		Leu	Gly	Ser	Ser 1300		Val	Ser	Ser	Arg 1305	Val	Ile	Val
45	Lys	Gly 1310		Arg	Asn		Ser 1315		Ser	Glu	Glu	Gly 1320		Val	Ile
50	Gly	Tyr 1325		Thr	Thr	Asp	Gly 1330		Leu	Leu	Gly	Ala 1335	Leu	Ser	Ile

	Gly	Asp 1340		Gly	Lys	Tyr	Arg 1345		Ile	Ile	Asn	Val 1350	Ala	Asp	Gly
5	Ser	Glu 1355	Ala	His	Asp	Ala	Val 1360	Thr	Val	Arg	Gln	Leu 1365	Gln	Asn	Ala
10	Ile	Gly 1370	Ala	Val	Ala	Thr	Thr 1375	Pro	Thr	Lys	Tyr	Tyr 1380		Ala	Asn
15	Ser	Thr 1385	Ala	Glu	Asp	Ser	Leu 1390	Ala	Val	Gly	Glu	Asp 1395	Ser	Leu	Ala
20	Met	Gly 1400	Ala	Lys	Thr	Ile	Val 1405	Asn	Gly	Asn	Ala	Gly 1410	Ile	Gly	Ile
•	Gly	Leu 1415	Asn	Thr	Leu	Val	Leu 1420	Ala	Asp	Ala	Ile	Asn 1425	Gly	Ile	Ala
25	Ile	Gly 1430	Ser	Asn	Ala	Arg	Ala 14 \$ 5	Asn	His	Ala	Asp	Ser 1440	Ile	Ala	Met
30	Gly	Asn 1445	Gly	Ser	Gln	Thr	Thr 1450	Arg	Gly	Ala	Gln	Thr 1455	Asn	Tyr	Thr
35	Ala	Tyr 1460	Asn	Met	Asp	Ala	Pro 1465	Gln	Asn	Ser	Val	Gly 1470	Glu	Phe	Ser
40	Val	Gly 1475	Ser	Glu	Asp	Gly	Gln 1480	Arg	Gln	Ilė	Thr	Asn 1485	Val	Ala	Ala
	Gly	Ser 1490	Ala	Asp	Thr	Asp	Ala 1495	Val	Asn	Val	Gly	Gln 1500	Leu	Lys	Val
45	Thr	Asp 1505	Ala	Gln	Val	Ser	Gln 1510	Asn	Thr	Gln	Ser	Ile 1515	Thr	Asn	Leu
50	Asn	Thr 1520	Gln	Val	Thr	Asn	Leu 1525	Asp	Thr	Arg	Val	Thr 1530	Asn	Ile	Glu

5	Asn	Gly 1535	Ile	Gly	Asp	Ile	Va 1540	Thr	Thr	Gly	Ser	Thr 1545	Lys	Tyr	Phe
	Lys	Thr 1550	Asn	Thr	Asp	Gly	Ala 1555	Asp	Ala	Asn	Ala	Gln 1560	Gly	Lys	Asp
10	Ser	Val 1565		Ile	Gly	Ser	Gly 1570	Ser	Ile	Ala	Ala	Ala 1575	Asp	Asn	Ser
15	Val	Ala 1580	Leu	Gly	Thr	Gly	Ser 1585	Val	Ala	Asp	Glu	Glu 1590	Asn	Thr	Ile
20	Ser	Val 1595	Gİy	Ser	Ser	Thr	Asn 1600	Gln	Arg	Arg	Ile	Thr 1605	Asn	Val	Ala
25	Ala	Gly 1610	Val	Asn	Ala	Thr	Asp 1615	Ala	Val	Asn	Val	Ser 1620	Gln	Leu	Lys
	Ser	Ser 1625	Glü	Дlа	Gly	Gly	Val 1630	Arg	Tyr	Asp	Thr	Lys 1635	Ala	Asp	Gly
30 ′	Ser	Ile 1640	Asp	Tyr	Ser	Asn	Ile 1645	Thr	Leu	Gly	Gly	Gly 1650	Asn	Ser	Gly
35	Thr	Thr 1655	Arg	Ile	Ser	Asn	Val 1660	Ser	Ala	Gly	Val	Asn 1665	Asņ	Asn	Asp
40	Ala	Val 1670	Asn	Tyr	Ala	Gln	Ļеи 1675	Lys	Gln	Ser	Val	Gln 1680	Glu	Thr	Lys
45	Gln	Tyr 1685		Asp ,	Gln	Arg	Met 1690	Val	Glu	Met	Asp	Asn 1695	Lys	Leu	Ser
	Lys	Thr 1700	Glu		Lys	Leu	Ser 1705	Gly	Ġly	Ile	Ala	Ser 1710	Ala	Met ,	Ala
50	Met	Thr	Gly	Leu	Pro	Gln	Ala	Tyr	Thr	Pro	Gly	Ala	Ser	Met	Ala

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Ser Ile Gly Gly Gly Thr Tyr Asn Gly Glu Ser Ala Val Ala Leu Ser Met Val Ser Ala Asn Gly Arg Trp Val Tyr Lys Leu Gly Val Gln Gly Ser Thr Asn Ser Gln Gly Glu Tyr Ser Ala Ala Leu Gly Ala Gly Ile Gln Trp <210> 35 <211> 227 <212> PRT <213> Escherichia coli <400> Met Asn Leu Lys Lys Thr Leu Leu Ser Val Leu Met Ile Leu Gln Cys Leu Leu Val Gly Cys Asp Tyr Ile Glu Lys Ala Ser Lys Val Asp Asp Leu Val Thr Gln Gln Glu Leu Gln Lys Ser Lys Ile Glu Ala Glu Lys Gln Gln Glu Leu Asp Lys Arg Lys Ile Glu His Phe Glu Lys Gln Gln Thr Thr Ile Ile Asn Ser Thr Lys Thr Leu Ala Gly Val Val

Lys Ala Val Lys Asn Lys Gln Asp Glu Phe Val Phe Thr Glu Phe Asn 50 85 90 95

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Pro Ala Gln Thr Gln Tyr Phe Ile Leu Asn Asn Gly Ser Val Gly Ala Gly Lys Ile Leu Ser Ile Asp Ala Val Glu Asn Gly Ser Val-Ile Arg Ile Ser Leu Val Asn Leu Leu Ser Val Pro Val Ser Asn Met Gly 5 Phe Tyr Ala Thr Trp Gly Gly Glu Lys Pro Thr Asp Ile Asn Ala Leu Ala Lys Trp Gln Gln Leu Leu Phe Ser Thr Ala Met Ash Ser Ser Leu Lys Leu Leu Pro Gly Gln Trp Gln Asp Ile Asn Leu Thr Leu Lys Gly Val Ser Pro Asn Asn Leu Lys Tyr Leu Lys Leu Ala Ile Asn Met Ala Asn Ile Gln Phe Asp Arg Leu Gln Pro Ala Glu Ser Pro Gln Arg Lys Asn Lys Lys <210> 36 <211> 1109 <212> PRT <213> Escherichia coli <400> 36

Met Lys Arg Val Val Arg Leu Leu Gly Val Gly Leu Leu Leu Val

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	1				5					10					15		
5	Val Ile	Leu	Leu		Ile	Leu	Phe	Val		Ala	Gln	Thr	Thr	Pro	Leu		
				20					25	•				30			
10	Ser Ala	Ala	Gln	Asp	Glu	His	Ala	Val	Trp	Leu	Arg	Leu	Leu	Ile	Thr		
	•	•	35		,			40					45	(*)			
15	Ile Phe	Val	Ile	Cys	Leu	Leu	Ser	Met	. Cys	Ile	Phe	Phe	Leu	Phe	Ser		
		50					55					60			٠		
20	Arg Asp	Gln	Asn	Glu	Ala	Ser	Thr	Ile	Ser	Leu	Tyr	Ala	Gln	Pro	Thr		
	65					70					75					80	
25	Ile Thr	Lys	Glu	Ile	Asn	Thr	Glu	Gln	Pro	Asn	Tyr	Ala	Ser	Leu	Leu		
					85					90					95		
30 ·	Ile Arg	Tyr	Leu	Arg	Asp	Arg	Tyr	Gly	Pro	Phe	Trp	Arg	Arg	Lys	Val		
		•		100					105					110			
35	Leu Pro	Leu	Leu	Val	Thr	Gly	Glu	Pro	Glu.	Gln	Ala	Glu	Ala	Ile	Ala		
			115					120					125				
40	Gly Ile	Ļeu	Thr	Gly	Gln	His	Trp	Leu	Glu	Gly	Asp	His	Thr	Val	Leu.		
		130					135				,	140				٠	
45	Tyr Ala	Gly	Gly	Arg	Pro	Thr	Ala	Glu	Pro	Asp	Val	Thr	Leu	Leu	Thr		
	145 160					150					155						
		4										•				•	

Leu Lys Lys Leu Arg Arg Ser Arg Pro Leu Asp Gly Ile Ile Trp Ala

165 170 175

Leu Thr Glu Glu Gln Ser Arg Gln Thr Ala Gln Leu Asp Lys Gly
5 Trp
180 185 190

Arg Gly Leu Ile Asn Gly Gly Lys Arg Leu Gly Phe Gln Ala Pro 10 Leu 195 200 205

Tyr Leu Trp Gln Val Cys Asp Asp Gly Asp Tyr Gln Thr Gly Arg
Pro
210 215 220

Leu Gln Ser Val Gly Cys Leu Leu Pro Glu Arg Cys Thr Pro Glu

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- 25 Leu Ala Val Met Leu Glu Ala Ala Ala Asp Gly Thr Gly His Val Ala 245 250 255
- 30 Ala Thr Asp Arg Tyr Arg Met Phe Ser Ala Ala Ser Gly Ser Tyr Pro
 260 265 270
- Cys Arg Ala Gly Tyr Cys Ser Leu Ala Asp Arg Pro Glu Thr Ala Ala 275 280 285
- 40 Gly Arg Arg Ile Phe Phe Pro Ala Pro Ala Arg Pro Asp Val Gln 290 295 300
- Pro Ala Ala Cys Arg Arg Ala Gly Gly Gln His Leu Met Gln Trp Leu
 305 310 315

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	Pro Ala	Ser	Pro	Val	Trp	Ala	Gly	Val	Thr	Val	Ile	Thr	Arg	Ala	Gly
					325					330					335
5	Arg	Trp	Val	Phe	Leu	Trp	Leu	Arg	Thr	Ala	Leu	Met	Ser	Ala	Val
	Cys			340					345					350	
10	77~7	Т О 17	\$7~ 1	T10	Trn	Glv	73.7 >	Glv	Met	Thr	Thr	Ser	Phe	Phe	Δla
	Asn	пец	355	116	ттЪ	OTY	FILA	360	1100	1111	7.1.1	501	365	1110	2114
15			330					•							
	Arg Asp	Ala	Leu	Val	Gln	Glu	Thr	Gly	Ile	Gln	Thr	Ala	Arg	Ala	Leu
		370					375			•		380			
20		Arg	Leu	Pro	Leu	Ala	Glu	Gln	Leu	Val	Ala	Leu	His	Thr	Leu
	Gln 385					390					395				
25	400														
	Gly Trp	Glu	Leu	Glu	Arg	Leu	Gln	Туг	Arg	Ile	Arg	Glu	Gly	Ala	Pro
30	. *				405					410				-	415
	_	Gln	Arg	Phe	Gly	Leu	Glu	Arg	Asn	Gln	Gln	Leu	Leu	Ala	Ala
2.5	Ala			420					425					430	
35	Dho	Dro.	Clv	ጥኒንድ	בומ	Gln	Δla	7 . 1 a	Asn	Ara	Leu	Val	Ara	Aso	Val
	Ala	FLO	435	тут	ALG	GTII	ALU	440	11011	,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,	110 a	Val	445	110p	var
40		•	,00												
	Val Pro	Asp	His	Leu	Gln	Gln	Gln	Leu	Asn	Ala	Phe		Ala	Leu	Pro
45		450			-		455		•			460			
		Ser	·Pro	Gln	Arg	Thr	Ala	Thr	Gly	Glu	Gln	Arg	Tyr	Lys	Gln
50	Leu 465 480					470					4 75				٠

	'	W O 200	US/US / C	23				119/	/370				rc1,	/EF 200	3/0039
		Ala	Leu	Leu	Met	Thr	Ser	Arg	Pro	Glu	Lys	Ala	Asp	Ala	Ala
5	Phe				485					490					495
		Ser	Thr	Thr	Leu	Met	Ala	Asp	Gly	Leu	Arg	T yr	Glu	Asn	Ile
10	Pro			500					505		·			510	
	Glu Thr	Gly	Val	Arg	Gln	Ser	Val	Leu	Pro	Ser	Leu	Leu	Thr	Phe	Trp
15	T11T		515					520,			٠		525		,
		Asn	Leu	Pro	Glu	His	Pro	Gln	Trp	Lys	Thr	Ser	Pro	Pro	Pro
20	Glu	530	·				535				ŕ	540			
		Thr	Gly	Ala	Val	Arg	Lys	Ile	Leu	Leu	Arg	Gln	Ile	Gly	Val
25	Arg 545 560					550			•		555				
		Ala	Glu	Asn	Thr	Leu	Tyr	Gln	Asn	Val	Leu	Gln	Gln	Val	Ser
30	Arg				565			•		57 <u>0</u>					575
			Ala	Asp	Met	Thr	Leu	Ala	Asp	Met	Thr	G l y	Asp	Thr	Leu
35	Thr			580					585					590	
			Leu	Phe	Ser	Thr	Glu	Gln	Thr	Val	Pro	Gly	Met	Phe	Thr
40	Arg		595	•				600					605		
			Trp	Glu	Gly	Gln	Val	Arg	Glu	Ala	Ile	GLu	Gln	Val	Val
45	Thr	610					615					620			

Ala Arg Arg Glu Glu Ile Asp Trp Val Leu Ser Asp Arg Gln Gln
Asp
50 625 630 635

	Thr Ser		Ala	Asp	Ile	Ser	Pro	Asp	Thr	Leu	Arg	Asn	Arg	Leu	Thr
5					645				٠	650					655
	Arg Ser	Tyr	Phe	Thr	Asp	Phe	Ala	Gly	Ser	Trp	Leu	Ala	Phe	Leu	Asn
10	DET			660	*				665					670	-
		His	Trp	Lys	Lys	Glu	Asp	Ser	Leu	Ser	Gly	Ile	Leu	Asp	Gln
15	Leu		675					680					685		
		Leu	Met	Ala	Āsp	Ala	Arg	Gln	Ser	Pro	Lẹu	Ile	Ala	Leu	Thr
20	Asp	690					695					700			
	Thr	Leu	Ala	Trp	Gln	Ala	Ala	Thr	Gly	Arg	Glu	Asn	Arg	Gly	Leu ·
25	Ser 705 720					710					715				
	70	0	T	70.71 -	T	a				_					
30	Lys	ser	теп	Ala	туs 725	ser	Ala	GIn	GLu	Leu 730	Phe	Asn	Gly	Lys	Glu 735.
							_			•					
35	Thr Leu	Pro	GIn	Gln	Ser	Arg	Glu	Gly		Asp	Val	Pro	.Val		Pro'
				740					745					750	
40	Asp Gly	Lys	Thr	Phe	Thr	Pro	Leu	Leu	Arg	Leu	Leu	Gly	Asp	Lys	Ala
	-		755					760				,	765		
45	Gly Thr	Gly	Asp	Ser	Gln	Leu	Ser	Leu	Gln	Thr	Tyr	Leu	Thr	Arg	Val
.3		770		•			775					780			
50	Arg Gln	Val	Arg	Leu	Lys	Leu	Gln	Gln	Val	Thr	Asn	Ala	Pro	Asp	Pro

785 790 795 800 ·

5 Glu Met Thr Gln Gln Leu Ala Gln Thr Val Leu Gln Gly Lys Thr Val 805 810 815

10 Asp Leu Thr Asp Thr Arg Asp Tyr Gly Arg Leu Ile Ala Ala Ser Leu 820 825 830

15 Gly Glu Glu Trp Ser Gly Phe Gly Gln Ala Leu Phe Val Arg Pro Val 835 840 845

20 Glu Gln Ser Trp Arg Gln Val Leu Thr Pro Ala Ala Asp Ser Leu Asn 850 855 860

25 Arg Gln Trp Gln Arg Ala Ile Val Ser His Trp Asn Gln Asp Phe Ala 865 870 875 880

30
Gly Arg Tyr Pro Phe Lys Ala Ser Gln Asn Asp Ala Ser Leu Pro Leu
885
890
895

Leu Ala Gln Tyr Leu Arg Asp Asp Gly Arg Ile Asn Leu Phe Ile Ala 900 905 910

40
Ala Asn Leu Ser Gly Val Leu Lys Arg Glu Gly Arg Tyr Trp Val
Ala
915
920
925

Asp Ala Met Asn Thr Gln Gly Leu Thr Val Asn Pro Asp Phe Ile Arg
930
935
940

122/370
122/3/0

5	Ala Gly 945 960		Asn	Arg	Leu	Arg 950	Asp	Val	Ala	Asp	Thr 955	Ala P	he Al	a Se	r .
10	Asp Val	Ala	Gly	Ile	His 965	Phe	Glu	Leu	Arg	Ala 970	Lys	Pro A	la Ar	g As	
15	Met Asn	Lys	Thr	His 980	Leu	Val	Ile	Asp	Gly 985	Gln	Glu :	Leu G.	lu Ty '99		е
20	Gln Gln	Lys	Glu 995	Arg	Trp	Gln		Phe 1000		Trp	Pro	Asp (Glu 1005	Gln :	Irp
25	,	Gly 1010)				101	5				n Ala 1020)	Glu	
30		1025 Gln 1040	Ala				103	0 Va				Ile 1035	Phe		
35	Val		Lys	Ala	Gln	Asp	104! Gly 1060	Le	u Pr	ó Le	u Asr	1050 n Tyr 1065	Leu	Leu	Arg
40	Val	Glu 1070	Gln	Gly	Lys	Gly	Pro 1075	Le 5	u Al	a Le	u Leu	ı Glu 1080		Lys	Asn
45	Phe	Arg 1085	Leu	Pro	Gly	Gl'n	Val 1090		e Le	u Th:	r Gly	/ Lys 1095		Met	Lys
50		1100					1105	j .			o Glu				
50	37	/ 3	1 <2.	T T >	Τ/Ω	<212	∠> Ŀ	'KT	<z13< td=""><td>> E'S</td><td>scher</td><td>ichia</td><td>coli</td><td>. <4</td><td>00></td></z13<>	> E'S	scher	ichia	coli	. <4	00>

	Met Met	Phe	Pro	Ile	Arg	Phe	Lys	Arg	Pro	Ala	Leu	Leu	Суз	Met	Ala	
5	1				5					10					15	
	Leu Asp	Thr	Val	Val	Leu	Ser	Gly	Cys	Gly	Leu	Ile	Gln	Lys	Val	Val	
10				20	•	. ,			25					30	•	
	Glu	Ser	Lys	Ser	Val	Ala	Ser	Ala	Val	Phe	Tyr	Lys	Gln	Ile	Lys	
15	Ile		35					40					45			
		His	Leu	Asp	Phe	Phe	Ser	Arg	Ser	Ala	Leu	Asn	Thr	Asp	Ala	
20	Glu	50					55					60				
20	Asp	Thr	Pro	Leu	Ser	Thr	Met	Val	His	Val	Trp	Gln	Leu	Lys	Thr	
25	Arg 65					70					,75					80
20	Glu	Asp	Phe	Asp	Lys	Ala	Asp	Tyr	Asp	Thr	Leu	Phe	Met	Gln	Glu	
	Glu	_		_	85			_		90					95	
30									,							
	Lys	Thr	Leu		Lys	Asp	Val	Leu		Lys	His	Thr	Val		Val	
35				100					105			·	•	110		
	Pro Gln	Glu	Gly	Thr	Ala	Ser	Leu	Asn	Val	Pro	Leu	Asp	Lys	Glu	Thr	
40			115					120					125			
		Val	Ala	Ile	Ile	Gly	Gln	Phe	Tyr	His	Pro	Asp	Glu	Lys	Ser	
15	Asp	130					135					140	,			·
45	Ser	Trp	Δrα	T,e11	Va1	Tla	T,ve	Ara	Asn	Glii	T,eu	Glu	ДΊа	Asn	Twe	
	Pro 145	P	1119	u	v U. J.	150	_y.	9			155	214			~ <u>,</u> 2	

50

								124/3′	7 0							
	Arg Lys	Ser	: Ile	: Glu	Leu 165	Met	Arg	ser Ser	: Asp	Leu - 170		, Leu	. Leu	Pro		
5					100					170					175)
	Asp	Lys														
10		•														
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15	Met Cys	Ile	Ser	Gly	Gly	Asn	Met	Leu	Lys	Glu	Trp	Met	Ile	Phe	Thr	
	1				5					10					15	
20	Ser Ile	Leu	Leu	Thr	Leu	Ala	Gly	Ala	Ser	Leu	Pro	Leu	Ser	Gly	Cys	
20	116			20					25					30		
25	Ser	Arg	Gly	Gln	Glu	Ser	Ile	Ser	Glu	Gly	Ala	Ala	Phe	Gly	Ala	
25	Gly		35					40					45			
	T 1-	T	77	~ 1	_	~ 7										
30	Leu		Arg	GLU	Pro	Gly		Thr	Lys	Lys	Ala	Asp	Thr	Lys	Asp	
		50	ř				55					60				
	Asn	Val	Pro	Pro	Pro	Val	Tyr	Gly	Pro	Pro	Gln	Val	Ile	Phe	Arq	
35	Ile 65					70					75				2	80
						•										
40	Asp Asn	Asp	Asn	Arg	Tyr	Phe	Thr	Leu	Glu	Asn	Tyr	Thr	His	Cys	Glu	
e e					85					90					95	
	Glv	Gln	Thr	Phe	ጥላታ	Aen	Δον	Tara	7\ 7\ ~	T	7\ ~	T7 -	172 -	77 - 77	T	
45	Ile	O T 11	T11T		т Л т	Asn	MSII	тўз		туѕ	ASN	тте	nls		туѕ	
				100.					105					110		

Leu Asp Ala Ser Gly Tyr Leu Phe Lys Gly Arg Leu Phe Trp Leu 50 Ser 115 120 120 125

	Thr His	Arg	Asp	Asp	Phe	Leu	Ala	Phe	Pro	Ala	Thr	Leu	Asn	Thr	Arg
5	urs	130					135				-	140			
	Val	Ser	Cys	Met	Gly		Asn	Lys	Gly	Cys		Asn	Ala	Val	.Ile
10	145 160			,	•	150					155			٠.	
15	Thr Tyr	Thr	Asp	Gly		Lys	Arg	Arg	Ser		Val	Pro	Tyr	Gly	
					165					170			,		175
20	Thr Met	Gln	Asn	Pro	Thr	Gly	Ala	Thr	Arg	Asp	Tyr	Asp	Met	Leu	Val
				180					185		•			190	
25	Asn Phe	Asp	Gly	Phe	Tyr	Leu	Ļeu	Arg	Tyr	Arg	Gly	Gly	Gln	Gly	Arg
25	11.0		195					200		•			205		
30	Ser Gly	Pro	Val	Ile	Leu	Arg	Trp	Ile	Leu	Ser	Thr	Glu	Asp	Ser	Ser
30	O _T	210					215	•			-	220			
35	Val Glu	Val	Arg	Ser	Glu	_	Ala	Tyr	Glu	Leu	*	Arg	Pro	Gly	Glu
	225 240		-6			230				a	235			•	
40	Val Pro	Pro	Ser	Thr	Gly	Phe	Tyr	Lys	Ile	Asp	Leu	Ser	Arg	Phe	Tyr
	110				245					250					255
45	Lys Val	Asn	Asn	Val	Met	Glu	Met	Gln	Cys	Asp	Arg	Thr	Leu	Glu	Pro
				260					265					270	
50	Gln	Pro	Ser 275	Glu	Ser	Lys	Ile	Gln 280							

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39 <211> 501 <212> PRT <213> Escherichia coli <400> <210> Met Glu His Val Ser Ile Lys Thr Leu Tyr His Leu Leu Cys Cys Met 5 . Leu Leu Phe Ile Ser Ala Met Cys Ala Leu Ala Gln Glu His Glu Pro Ile Gly Ala Gln Asp Glu Arg Leu Ser Thr Leu Ile His Gln Arg Met Gln Glu Ala Lys Val Pro Ala Leu Ser Val Ser Val Thr Ile Lys Gly 55 ' Val Arg Gln Arg Phe Val Tyr Gly Val Ala Asp Val Ala Ser Gln Lys Ala Asn Thr Leu Asp Thr Val Tyr Glu Leu Gly Ser Met Ser Lys Ala Phe Thr Gly Leu Val Val Gln Ile Leu Ile Gln Glu Gly Arg Leu Arg Gln Gly Asp Asp Ile Ile Thr Tyr Leu Pro Glu Met Arg Leu Asn Tyr Gln Gly Lys Pro Ala Ser Leu Thr Val Ala Asp Phe Leu Tyr His Thr

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		127/370														
	_	Ser Pro 145 160	5	y Le	u Pro	⊃ Ph∈	Sei 150		. Leu	ı Ala	a Arg	155		u As:	n Pro	o Met
	5	Gl _y Ala	7 Se:	r Ala	a Val	_ Ala	Glr	ı Gln	Leu	l Arg	(Asn	Glu	ı Ası	n Lei	ı Let	ı Phe
	10					165	1				170					175
		Pro Gly	Gly	y Ala		Phe	Ser	Tyr	Ala			Asn	туг	a As <u>r</u>	o Val	Leu
	15			*	180					185					190	
		Ala Ala	Val	L Il∈	e Glu	Asn	Val	Thr	Gly	Lys	Thr	Phe	Thr	Glu	ı Val	Ile
	20			195					200					205	j	•
		Glu Lys	Arç	g Leu	Thr	Gln	Pro	Leu	Gly	Met	Ser	Ala	Thr	Val	Ala	Val
	25		210)				215				•	220			
		Pne	Asp	Glu	Ile	Ile	Val	Asn	Lys	Ala	Ser	Gly	Tyr	Lys	Leu	Gly
	30	225 240					230	,			•	235				
		Gly	Lys	Pro	Val	Leu	Phe	His	Ala	Pro	Leu	Ala	Arg	Asn	His	Val
	35	Pro				245	,				250		•			255
		Ala Asp	Ala	Tyr	Ile	His	Ser	Thr	Leu	Pro	Asp	Met	Glu	Ile	Trp	Ile
	40				260					265					270	
		Ala Met	Trp	Leu	His	Arg	Lys	Ala	Leu	Pro	Ala	Thr	Leu	Arg	Glu	Ala
45			275					280				•	285			
		Ser Asn	Asn	Ser	Trp	Arg	Gly	Asn	Ser	Asp	Val	Pro	Leu	Ala	Ala	Asp
4	50		290					295					300			

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5	Arg Pro 305 320	Ile	Leu	Tyr	Ala	Ser 310	Gly	Trp	Phė	Ile	Asp 315	Ģln	Asn	Gln	Gly
10	Tyr Ala	Ile	Ser	His	Gly 325	Gly	Gln	Asn	Pro	Asn 330	Phe	Ser	Ser	Cys	Ile 335
15	Leu Asn	Arg [°]	Pro	Asp 340	Gln	Gln	Ile	Gly	Ile 345	Val	Ala	Leu	Ala	Asn 350	Met
20	Ser Arg	Asn	Leu 355	Ile	Leu	Gln	Leu	Cys 360	Ala	Asp	Ile	Asp	Asn 365	Tyr	Leu
25	Ile Asp	Gly 370	Lys	Tyr	Ala	Asp	Gly 375	Ala	Gly	Asp	Ala	Ile 380	Thr	Ala	Thr
30	Thr Val 385 400	Leu	Phe	Val	Tyr	Leu 390	Thr	Leu	Leu	Leu	Cys 395	Phe	Trp	Gly	Ala
35	Val Gly	Val	Val	Arg	Gly 405		Phe	Arg	Val	Tyr 410	Arg	Ala	Thr	Ala	His 415
40	Pro Ile	Gly	Lys	Gln 420	Gln	Arg	Leu	Arg	Leu 425	Arg	Val	Arg	Asp	Tyr 430	Ile
45	Ala Pro	Leu	Ala 435	Val	Pro	Gly	Leu	Val 440		Ala	Met	Leu	Tyr 445	Val	Ala
50	Gly Gly		Leu	Ser	Pro	Gly	Leu 455	Asp	Trp	Arg	Phe	Ile	Leu	Val	Trp

5	Pro Ser Ser Phe 465 480	Val Leu	Ala Ile	Pro Phe	Gly Ile 3	Ile Leu Leu	Ala
10	Val Leu Thr Lys	Leu Asn	His Gln	Ile Lys	Arg Ile 1	Leu Leu His	Asn 495
15	Glu Trp Asp	Asp Glu					
20	<210> 40 <2	211> 68	2 <212>	PRT <213	3> Esche:	richia coli	<400>
20	Met Lys Asn Ser 1	Lys Tyr	Ile Ile	Ala Pro	Gly Ile 7	Ala Val Met	Cys 15
25	Ala Val Ile Thr	Ser Ser	Gly Tyr	Ala Ser	Ser Asp	Lys Lys Glu 30	Asp
30	Leu Val Val Pro 35	Thr Ala	Ser Gly	Phe Thr	Gln Gln	Leu Arg Asn 45	Ala
35	Ala Ser Val Val .	Ser Val	Ile Thr	Ser Glu		Gln Lys Lys 60	Pro
40	Ser Asp Leu Thr 65	Val Asp	Ala Val	Lys Asp	Val Glu	Gly Ile Ser	Ile 80
45	Gly Gly Asn Asp	Glu Lys	Pro Asp	Ile Ser	Ile Arg	Gly Leu Ser	,Gly 95

V	WO 200	05/0978	323				130,	/370				PC	Г/ЕР20	05/0039	72
Tyr Ser	Thr	Leu	Ile	Leu	Val	Asp	Gly	Arg	Arg	Gln	Ser	Gly	Arg	Glu	-
per			100					105					110		

Arg Pro Asn Gly Ser Gly Gly Phe Glu Ala Gly Phe Ile Pro Pro Val

115
120
125

10

15

20

25

30

35

40

45

Glu Ala Ile Glu Arg Ile Glu Val Ile Arg Gly Pro Met Ser Ser Leu 130 135 140

Tyr Gly Ser Asp Ala Ile Gly Gly Val Ile Asn Ile Ile Thr Lys Pro 145 150 155

Val Asn Asn Gln Thr Trp Asp Gly Val Leu Gly Leu Gly Gly Ile Ile 165 170 170

Gln Glu His Gly Lys Phe Gly Asn Ser Thr Thr Asn Asp Phe Tyr Leu
180 185 190

Ser Gly Pro Leu Ile Lys Asp Lys Leu Gly Leu Gln Leu Tyr Gly Gly 195 200 205

Met Asn Tyr Arg Lys Glu Asp Ser Ile Ser Gln Gly Thr Pro Ala Lys 210 215 220

Asp Asn Lys Asn Ile Thr Ala Thr Leu Gln Phe Thr Pro Thr Glu Ser
225 230 235
240

Gln Lys Phe Val Phe Glu Tyr Gly Lys Asn Asn Gln Val His Thr Leu 50 245 250 255

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Thr Pro Gly Glu Ser Leu Asp Ala Trp Thr Met Arg Gly Asn Leu Lys Gln Pro Asn Ser Lys Arg Glu Thr His Asn Ser Arg Ser His Trp Val Ala Ala Trp Asn Ala Gln Gly Glu Ile Leu His Pro Glu Ile Ala Val Tyr Gln Glu Lys Val Ile Arg Glu Val Lys Ser Gly Lys Lys Asp Lys Tyr Asn His Trp Asp Leu Asn Tyr Glu Ser Arg Lys Pro Glu Ile Thr Asn Thr Ile Ile Asp Ala Lys Val Thr Ala Phe Leu Pro Glu Asn Val 345 . 30 : Leu Thr Ile Gly Gly Gln Phe Gln His Ala Glu Leu Arg Asp Asp Ser Ala Thr Gly Lys Lys Thr Thr Glu Thr Gln Ser Val Ser Ile Lys Gln Lys Ala Val Phe Ile Glu Asn Glu Tyr Ala Ala Thr Asp Ser Leu Ala Leu Thr Gly Gly Leu Arg Leu Asp Asn His Glu Ile Tyr Gly Ser Tyr

															•
	Trp Thr	Asn	Pro	Arg	Leu	Tyr	Ala	Val	Tyr	Asn	Leu	Thr	Asp	Asn	Leu
5				420					425					430	
	Leu Glu	Lys	Gly	Gly	Ile	Ala	Lys	Ala	Phe	Arg	Ala	Pro	Ser	Ile	Arg
10	GIU	•	435					440					445		
		Ser	Pro	Gly	Phẹ	Gly	Thr	Leu	Thr	Gln	Gly	Gly	Ala	Ser	Ile
15	Met	450				٠	455				-	460·			
	Tyr	Gly	Asn	Arg	Asp	Leu	Lys	Pro	Ģlu	Thr	Ser	Val	Thr	Glu	Glu
20	11e 465 480					470					475				
	400											•			
25	Gly Leu	Ile	Ile	Tyr		Asn	Asp	Ser	Gly	Phe	Ser	Ala	Ser	Ala	
					485	,				490					495
30	Phe Thr	Asn	Thr	Asp	Phe	Lys	Asn	Lys	Leu	Thr	Ser	Tyr	Asp	Ile	Gly
				500					505					510	,
35	Lys Gly	Asp	Pro	Val	Thr	Gly	Leu	Asn	Thr	Phe	Ile	Tyr	Asp	Asn	Val
	GIY		515					520					525		
40		Ala	Asn	Ile	Arg	Gly	Val	Glu	Leu	Ala	Thr	Gln	Ile	Pro	Val
40	Tyr	530	-				535					540			
	_	Lys	Trp	His	.Val	Ser	Ala	Asn	Tyr	Thr	Phe	Thr	Asp	Ser	Arg
45	Arg 545 560					550		•		·	555				
	200					ē						,			
50	Lys Pro	Ser	Asp	Asp	Glu	Ser	Leu	Asn	Gly	Lys	Ser	Leu	Lys	Gly	Glu

	W	V O 2 00	5/0978	23				133/3	70				PCT/I	E P2 005	/003972
					565					570					575
5	Leu Asp	Glu	Arg	Thr 580	Pro	Arg	His	Ala	Ala 585	Asn	Ala	Lys	Leu	Glu 590	Trp
10	Tyr Lys	Thr	Gln 595	Asp	Ile	Thr	Phe	Tyr 600	Ser	Ser	Leu	Asn	Tyr 605	Thr	Gly
15	Gln Arg	Ile 610	Trp	Ala	Ala	Gln	Arg 615	Asn	Gly	Ala	Lys	Val	Pro	Arg	Val
20	Asn Pro 625 640	Gly	Phe	Thr	Ser	Met 630	Asp	Ile	Gly	Leu	Asn 635	Tyr	Gln	Ile	Leu
25	Asp Ser	Thr	Leu	Ile	Asn 645	Phe	Ala	Val	Leu	Asn 650	Val	Thr	Asp	Arg	Lys 655
30	Glu Arg	Asp	Ile	Asp 660	Thr	Ile	Asp	Gly	Asn	Trp	Gln	Val	Asp	Glu 670	Gly
35	Arg	Tyr	Trp 675	Ala	Asn	Val	Arg	Val 680	Ser	Phe					
40	<210 41)> 4	11 <2	211>	164	1 <23	L2> .	PRT	<213	3> E	Esche	erich	nia c	coli	<400>
45	Met Ile 1	Gly	Phe	Arg	Lys 5	Thr	Ile	Ile	Thr	Ser	Val	Gly	Leu	Ile	Phe 15
	Ser Asn	Phe	Ser	Phe	Val	Ala	Lys	Cys	Ser 25	Gln	Leu	Lys ·	Asn	Leu 30	Asn

	W	O 200	5/09782	23				134/3′	70				PCT/I	E P2 005	5/00 3 97	2
		Ser	Val	Met	Leu	Cys	Gly	Lys	Val	Ser	Asn	Asn	Ile	Leu	Asp	
	Asp		35					40					45			
5		Gly	Gly	Tyr	Lys	Glu	Arg	Asn	Ile	Leu	Met	Leu	Arg	Ala	Ile	
	Lys	50					55					60			-	
10		•			•					•						
	Lys Phe	Ile	Ile	Ile	Met	Thr	Ile	Val	Asn	Ile	Ile	Phe	Phe	Tyr	Ser	
	65					70					75				,	8 0
15	Gln Gly	Ser	Thr	Ala	Asp	Glu	Met	Val	Leu	Ile	Lys	Lys	Tyr	Gly	Phe	
	O±y				85					90					95	
20		Glu	Arg	Asp	Ile	Lys	Gl'y	Arg	Pro	Leu	Ile	Tyr	Pro	Ile	Glu	
	Asn			100					105					110		
25	ጥህድ	Asn	Glu	Cvs	T.vs	Lys	T.vs	Cvs	Δsn	Ніа	Met.	Δsn	Ψαν	Tle	70.1 =	
	Asp	мэр		Cys	шуз	د برند	пуз	_	ASII	1112	Mec	ASII		116	ALA	
20			115					120					125			
30																

Val Asn Ala Gln Leu Ala Met Ser Lys Lys Asn Asn Arg Ile Phe Ala

Asn Ile Thr Phe Thr Asn Asn Ser Ser Thr Thr Tyr Phe Phe Leu Asn

Ile Ile Tyr Leu

<210> 42 <211> 218 <212> PRT <213> Escherichia coli <400> Met Asn Gln Ile Lys Asp Asn Lys Val Ile Met Lys Ile Lys Asn Leu

	.Ile Tyr	Ser	Val	Ile	Leu	Leu	Ser	Gly	Gly	Ile	Met	Gly	Thr	Gly	Leu		
5	. T Y L			20					25					30			
	Ser	Ser	Asp	Asn	His	Gln	Lys	Ile	Arg	Ser	Arg	Phe	Asn	Ile	Gln		
10	Glu		35					40					45				
	a		Q	70 T -	T7 -	T	mb a	7) = 20	C1	77~ T	T 011	Clar	Dho	Sor	7\		
15	Ser	Tyr 50	Cys	Ala	TTE	гу	55	ASII	Gly	vai	пеп	60 GT 3	FIIC	per	ASII		
13		50					00		~						•		
	Lys Ser	Asp	Val	Leu	Arg	Glu	Asn	Gly	Asp	Ser	Thr	Gly	Thr	Thr	Ser		
20	65					70					75					80	
		Thr	Asn	Ala	Met	Met	Leu	Met	Glu	Asn	Gly	Glu	Asn	Glu	Ile		
25	Ser				85		•			90					95		•
	T.e.11	Glu	Tle	Glv	Ala	Leu	Ara	Tro	Phe	Ser	Asp	Lvs	Pro	Ala	Ser		
30	Thr	Olu	110	100			5		105		_			110			
						•											
	Glu Leu	Glu		Gly	His	Phe	Ser		Lys	Ala	Gly	Cys		Leu	Asp		•
35			115					120					125				
	Val Val	Arg	Phe	Val	Lys	Gln	Glu	Glu	Thr	Ile	Leu	Ser	Ser	Ile	Lys		
40	val	130			-		135					140					
	Thr	Ile	Asn	Gln	Gln	Gly	Ile	Pro	Glu	Ala	Gln	Pro	Asp	Ser	Met		
45	His 145					150					155	•					
	160			-													
50			Ile	Arg	Lys	Glu	Ile	Leu	Ala	Glu	Gln	Ala	Glu	Pro	Gly		
50	Phe				165					170	ı				175		

5	Ile Asp Lys	Pro	Asp	Tyr	Phe	Asņ	Glu	Thr 185	Tyr	Phe	Pro	Lys	Gly 190	Met	
10	Val Tyr Gly	Gln 195	Phe	Thr	Gln	Lys	Val 200	Ser	Val	Ala	Gly	Leu 205	Pro	Asp	
15	Pro Gly	Arg	Ser	Thr	Pro	Phe 215	Thr	Gly	Ala						
		43 <2 43	211>	273	32 <2	212>	PR	r <23	13>	Escl	heri	chia _.	col	Ĺ	,
20	Met His Leu 1	Gln	Pro	Pro 5	Val	Arg	Phe	Thr	Tyr 10	(Arg	Leu	Leu	Ser	Tyr 15	
25	Val Ser Ala	Ala	Ile 20	Ile	Ala	Gly	Gln	Pro 25	Leu	Leu	Pro	Ala	Val	Gly	
30	Val Ile Gly	Thr	Pro	Gln	Asn	Gly	Ala 40	Gly	Met	Asp	Lys	Ala 45	Ala	Asn	
35	Val Pro His 50	Val	Val	Asn	Ile	Ala 55	Thr	Pro	Asn	Gly	Ala 60	Gly	Ile	Ser	
40	Asn Arg Asn 65	Phe	Thr	Asp	Tyr 70	Asn	Val	Gly	Lys	Glu 75	Gly	Leu	Ile	Leu	80
45	Asn Ala Gln	Thr	Gly	Lys 85	Leu	Asn	Pro	Thr	Gln 90	Leu	Gly	Gly	Leu	Ile 95	
50	Asn Asn Asn	Pro	Asn	Leu	Lys	Ala	Gly	Gly	Glu	Ala	Lys	Gly	Ile	Ile	

Asn .

100 105 110

Glu Val Thr Gly Gly Lys Arg Ser Leu Leu Gln Gly Tyr Thr Glu 5 Val 115 120 125

Ala Gly Lys Ala Ala Asn Val Met Val Ala Asn Pro Tyr Gly Ile 10 Thr 130 135 140

Cys Asp Gly Cys Gly Phe Ile Asn Thr Pro His Ala Thr Leu Thr

Thr

145
160

20 Gly Lys Pro Val Met Asn Ala Asp Gly Ser Leu Gln Ala Leu Glu Val
165 170 175

25 Thr Glu Gly Ser Ile Thr Ile Asn Gly Ala Gly Leu Asp Gly Thr Arg
180 185 190

30 Ser Asp Ala Val Ser Ile Ile Ala Arg Ala Thr Glu Val Asn Ala Ala 195 200 205

35 Leu His Ala Lys Asp Leu Thr Val Thr Ala Gly Ala Asn Arg Val Thr 210 215 220

40 Ala Asp Gly Arg Val Arg Ala Leu Lys Gly Glu Gly Asp Val Pro Lys 225 230 235

Val Ala Val Asp Thr Gly Ala Leu Gly Gly Met Tyr Ala Arg Arg
Ile
245
250
255

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	His Leu	Leu	Thr	Ser	Thr	Glu	Ser	Gly	Val	Gly	Val	Asn	Leu	Gly	As'n
	200			260			٧.		265					270	
5	Tyr	Ala	Arg	Asp	Gly	Asp	Ile	Thr	Leu	Asp	Ala	Ser	Glv	Ara	Leu
	Thr		275		-	-		280		- 1			285	9	
10															
,	Val _. Gly		Asn	Ser	Leu	Ala	Thr	Gly	Ala	Val	Thr	Ala	Lys	Gly	Gln
		290					295	-				300			
15	Val	Thr	Leu	Thr	Gly	Asp	His	Lys	Ala	Gly	Gly	Asn	Leu	Ser	Val
	Ser					310					315				
20	320														
		Arg	Arg	Asp	Ile	Val	Leu	Ser	Asn	Gly	Thr	Leu	Asn	Ser	Asp
25	Lys				325					330					335
23	7) ~~	Т о	0	Т	m1	70 T -	C 1	~ 1	~	7					
	Lys	ьeu	ser		rnr	Ата	GIY	GTÀ		TTE	Thr	Gln	Gln		Glu
30				340				÷	345	•				350	
	Leu Gln	Thr	Ala	Gly	Arg	Asp	Val	Thr	Leu	Ala	Ala	Lys	Asn	Ile	Thr
35	GIII		355					360					365		
55	Asp	Thr	Ala	Ser	Gln	Tle	Asn	Δla	7\] a	Δκα	7) 52 70	Ile	™ 1	mb v	
-	Ala	370	I I I C	001	CIII	110	375	11.L.C	пта	Arg	Ash	380	val	T11T	Val
40							373					300			
	Ser Leu	Asp	Thr	Leu	Thr	Thr	Gln	Gly	Gln	Ile	Thr	Ala	Gly	Gln	Asn
45	385 400					390					395				
	Thr Lys	Ala	Ser	Ala	Thr	Thr	Leu	Thr	Gln	Asp	Gly	Ile	Leu	Leu	Ala
50					405	-				410	. •				415

•														-	
	Ser Val	His	Ala	Gly	Leu	Asn ·	Ala	Gly	Thr	Leu	Asn	Asn	Ser	Gly	Ala
5	val			420					425					430	
J	C1	C1	71.5	ШЬх	Tou	Th ∞	Т олл	C1	202	መb ኤ	™h r	T. 011	202	7) 0 22	C 0 20
	Gly	сту	Ala	THE	ьеи	Tift	теп	_	ser	TIIT	TIIT	ьеи	•	ASII	ser
10			435					440					445		
•		Leu	Leu	Ser	Gly	Gly.	Pro	Leu	Thr	Met	Asn	Thr	Arg	Asp	Phe
	Thr	450					455					460			
15															
	Gln Ser	Ser	Gly	Arg	Thr	Gly	Ala	Lys	Gly	Lys	Val	Asp	Ile	Met	Ala
20	465 480					470			ot .		475		ı		
	Gly Leu	Lys	Leu	Thr	Ser	Thr	Gly	Leu	Leu	Val	Thr	Met	His	Leu	Val
25	пец				485					490					495
	T	71 7	Gln	7, 00	V -1	ΠЬх	Cln	7\ c x	Clu	Va l	Lou	Sor	Clv	Clv	Tuc
20	Gly	Ala	GTII		vaı	7117	GIII	ASII		vai	шеи	Der	GTĀ		ту
30			·	500				,	505	ė		,		510	
		Thr	Val	Ser	Ala	Thr	Ser	Ser	Gly	Lys	Lys	Ser	Val	Thr	His
35	Ser		515		Ŷ			520					525		
	Asp Glu	Ala	Ala	Met	Thr	Leu	Asn	Val	Thr	Thr	Val	Ala	Leu	Asp	Gly
40		530					535			,		540			
	Thr	Ser	Ala	Gly	Asp	Thr	Leu	Arg	Val	Gln	Ala	Asp	Lys	Leu	Ser
45	Thr 545			_	_	550					555				
	560					- 1									
	71 ~	70.7 ~	Gly	7\ 1 -	Glin	T.A.	Gln	Sar	Cl to	T.ve	Zen	T, <u></u>	Sar	Tle	Aen
50	Ala	ATG	ату	ΔIA		пец	OTII	DGT	OTY	БуS 570		шcu	DOL		575
					565					570					515

												•			
	Arg Val	Asp	Ala	Arg	Leu	Ala	Gly	Thr	Gln	Ala	Ala	Gln	Gln	Thr	Met
5				580					585			•		590	
	Val Pro	Asn	Ala	Ser	Glu	Lyś	Leu	Thr	His	Ser	Gly	Lys	Ser	Ser	Ala
10			595					600					605		
	Ser Val		Ser	Leu	Ser	Ala	Pro	Glu	Leu	Thr	Ser	Ser	Gly	Val	Leu
15	, ,	610			•		615					620			
	Gly Leu	Ser	Ala	Leu	Asn	Thr	Gln	Ser	Gln	Thr	Leu	Thr	Asn	Ser	Gly
20	625 640					630			•	٠	635				-
~ ~		Gln	Gly	Glu	Ala	Ser	Leu	Thr	Val	Asn	Thr	Gln	Arg	Leu	Asp
25	Asn	•			645					650				-	655
		Gln	Asn	Gly	Thr	Leu	Tyr	Ser	Ala	Ala	Asp	Leu	Thr	Leu	Asp ·
30	Ile			660					665					670	
35	Pro Met	Asp	Ile	Arg	Asn	Ser	Gly	Leu	Ile	Thr	Gly	Asp	Asn	Gly	Leu
30	1100	•	675					680					685		
40	Leu Thr	Asn	Ala	Val	Ser	Leu	Ser	Asn	Pro	Gly	Lys	Ile	Ile	Ala	Asp
	1111	690					695					700.			
15	Leu	Ser	Val	Arg	Ala	Thr	Thr	Leu	Asp	Gly	Asp	Gly	Leu	Leu	Gln
45	Gly 705 720					710					715				
				•											

50 Ala Gly Ala Leu Ala Gly Asp Thr Leu Ser Gln Gly Ser His

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725 730 735

- Gly Arg Trp Leu Thr Ala Asp Asp Leu Ser Leu Arg Gly Lys Thr 5 Leu 740 745 750
- Asn Thr Ala Gly Thr Thr Gln Gly Gln Asn Ile Thr Val Gln Ala
 10 Asp
 755 760 765
- Ser Ala Thr Gly Gln Leu Thr Ser Thr Gly Asp Ile Met Ser Gln

 Gly
 785 790 795
- 25 Asp Thr Thr Leu Lys Ala Ala Thr Thr Asp Asn Arg Gly Ser Leu Leu 805 810 815
- 30 Ser Ala Gly Thr Leu Ser Leu Asp Gly Asn Ser Leu Asp Asn Arg Gly 820 825 830
- 35 Thr Val Gln Gly Asn His Val Thr Ile Arg Gln Asn Ser Val Thr Asn 835 840 845
- 40 Ser Gly Thr Leu Thr Gly Ile Ala Ala Leu Thr Leu Ala Ala Arg Met 850 855 860
- 45 Ala Ser Pro Gln Pro Ala Leu Met Asn Asn Gly Gly Ser Leu Leu Thr 865 870 875

	WO 2005/097823								370				PCT	/EP200	05/003972
		Gly	Asp	Leu	Thr	Ile	Thr	Ala	Gly	Ser	Ile	Thr	Ser	Ser	Gly
	His				885				•	890					895
	Trp Ser	Gln	Gly	Lys 900	Arg	Val	Leu	Ile		Ala	Asp	Ser	Leu		Asn
10	Gly Glu	Ala	Ile 915		Ala	Ala	Asp	Ser 920	905 Leu	Thr	Ala	Arg		910 Thr	Gly
15	Leu Ala	Val		Thr	Ala	Gly	Ser 935		Val	Thr	Ser	Asn	925 Gly	Glu	Met
20	Leu Asn 945 960		Ala	Leu	Asn	Leu 950	Ser	Asn	Ser	Gly ·	Gln 955		Ile	Ala	Lys
30	Leu Gly	Thr	Leu	Lys	Ala 965	Asn	Ser	Leu	Thr	Ser 970	Ala	Gly	Asp	Ile	Thr 975
35	Val Ala	Asp	Thr	Leu 980	Thr	Leu	Thr	Val	Asn 985	Gln	Thr	Leu	Asn	Asn 990	Gln
40	Asn Ser	Gly	Lys 995	Leu	Leù	Ser	Ala	Gly 1000		l Lei	ı Thi	: Lei	100		la Asp
	Val	Thr 1010		n Asp	o Gly	/ Glr	101		ln Gl	ly. As	sn Va		nr 1 020	Thr I	Ile Thr
45	Ala	Gly 1025		. Let	ı Thi	r Asr	1 Gly		.y Hi	is L∈	eu Gl		Ly ()35	Slu :	Thr Leu

Thr Leu Thr Ala Ser Gly Gly Val Asn Asn Arg Ser Gly Gly Val

5	Leu	Met 1055	Ser	Arg	Asn	Ala	Leu 1060		Val	Ser	Thr	Ala 1065		Leu	Ser
10		Gln 1070	Ser	Thr	Ile	Gln	Gly 1075	Gly	Gly	Gly	Val	Ser 1080		Asn	Ala
10 .		Asp 1085	Arg	Leu	Gln	Asn	Asp 1090	Gly	Lys	Ile	Leu	Ser 1095		Ser	Asn
15	Leu	Thr 1100	Leu	Thr	Ala	Gln	Val 1105	Leu	Ala	Asn	Thr	Gly 1110		Gly	Leu
20	Val	Gln 1115	Ala	Ala	Thr	Leu	Leu 1120	Leu	Asp	Val	Val	Asn 1125		Val	Asn
25	Gly	Gly 1130	Arg	Val	Lẹu	Ala	Thr 1135	Gly	Ser	Asp	Val	Lys 1140		Thr	Thr
	Leu	Asn 1145	Asn	Thr	Gly	Thr	Leu 1150	Gln	Gly	Ala	Thr	Leu _. 1155	Val	Asn	Tyr
30	His	Thr 1160	Phe	Ser	Ser	Gly	Thr 1165	Leu	Leu	Gly	Thr	Ser 1170	Gly	Leu	Gly
35	Val	Lys 1175	Gly	Ser	Ser	Leu	Leu 1180	Gln	Asn	Gly	Thr	Gly 1185	Arg	Leu	Tyr
40	Ser	Ala 1190	Gly	Asn	Leu	Leu	Leu 1195	Asp	Ala	Gln	Asp	Phe 1200	Ser	Gly	Gln
45	Gly	Gln 1205	Val	Val	Ala	Thr	Gly 1210	Asp	Val	Thr	Leu	Lys 1215	Leu ·	Ile	Ala
	Ala	Leu 1220	Thr	Asn	His	Gly	Thr 1225	Leu	Ala	Ala	Gly	Lys 1230	Thr	Leu	Ser
50	Val	Thr	Ser	Gln	Asn	Ala	Ile	Thr	Asn	Gly	Gly	Val	Met	Gln	Gly

1235	1240	1245 .

				•											
5	Asp	Ala 1250	Met	Val	Leu	Gly	Ala 1255	Gly	Glu	Ala	Phe	Thr 1260	Asn	Asn	Gly
10	Leu	Thr 1265		Gly	Lys	Gly	Asn 1270	Ser	Val	Phe	Ser	Ala 1275	Gln	Arg	Leu
	Phe	Leu 1280	Asn	Ala	Pro	Gly	Ser 1285	Leu	Gln	Gly	Gly	Gly 1290		Val	Ser
15	Leu	Asn 1295	Ser	Arg	Ser	Asp	Ile 1300	Thr	Ile	Ser	Gly	Phe 1305	Thr	Gly	Thr
20	Ala	Gly 1310	Ser	Leu	Thr	Met	Asn 1315	Val	Ala	Gly	Thr	Leu 1320	Leu	Asn	Ser
25	Ala	Leu 1325	Ile	Tyr	Ala	Gly	Asn 1330	Asn	Leu	Lys	Leu	Phe 1335	Thr	Asp	Arg
30	Leu	His 1340	Asn	Gln	His	Gly	Asp 1345	Ile	Leu	Ala	Gly	Asn 1350	Ser	Leu	Trp
	Val	Gln 1355	Lys	Asp	Ala	Ser	Gly 1360	Gly	Ala	Asn	Thr	Glu 1365	Ile	Ile	Asn
35	Asn	Ser 1370	Gly	Asn	Ile	Glu	Thr 1375	His	Gln	Gly	Asp	Ile 1380	Val	Val	Arg
40	Thr	Gly 1385	His	Leu	Leu	Asn	Gln 1390	Arg	Glu	Gly	Phe	Ser 1395	Ala	Thr	Thr
45	·Thr	Thr 1400	_	Thr	Asn	Pro	Ser 1405		Ile	Gln ·	Gly	Met 1410	Gly	Asn	Ala
50	Leu	Val 1415	Asp	Ile	Pro	Leu	Ser 1420	Leu	Leu	Pro	Asp	Gly 1425	Ser	Tyr	Gly

		WO 2 00	5/0978	23				145/37	0		•	P	CT/EP	2005/00	03972
	Tyr	Phe 1430		Arg	Glu	Val	Glu 1435			His	Gly	Thr 1440	Pro	Cys	Asn
5	Gly	His 1445		Ala	Cys.	Asn	Ile 1450	Thr	Met	Asp	Thr	Leu 1455	, Tyr	Tyr	Tyr
10	Ala	Pro 1460	Phe	Ala	Asp	Ser	Ala 1465	Thr	Gln	Arg	Phe ·	Leu 1470	Ser	Ser	Gln
.15	Asn	Ile 1475		Thr	Val	Thr	Gly 1480	Ala	Asp	Asn	Pro	Ala 1485			Ile '
20	Ala	Ser 1490	Gly	Arg	Asn	Leu	Ser 1495	Ala	Glu	Ala	Glu	Arg 1500	Leu	Glu	Asn
20	Arg	Ala 1505		Phe	Ile	Leu	Ala 1510	Asn	Gly	Asp	Ile	Ala 1515	Leu	Ser	Gly
25	Arg	Glu 1520	Leu	Ser	Asn	Gln	Ser 1525	Trp	Gln	Thr	Gly	Thr 1530	Glu	As'n	Glu
30	Tyr	Leu 1535	Val	Tyr	Arg	Tyr	Asp 1540	Pro	Lys	Thr	Phe	Tyr 1545	Gly	Ser	Tyr
35	Ala	Thr 1550	Gly	Ser	Leu	Asp	Lys 1555	Leu	Pro	Leu	Leu	Ser 1560	Pro	Glu	Phe
		Asn 1565	Asn	Thr	Įlе	Arg	Phe 1570	Ser	Leu	Asp	Gly	Arg 1575	Glu	Lys	Asp
40	Туг	Thr 1580	Pro	Gly	Lys	Thr	Tyr 1585	Tyr	Ser	Val	Ile	Gln 1590	Ala	Gly	Gly
45	Asp	Val 1595	Lys	Thr	Arg		Thr 1600	Ser	Ser	Ile	Asn	Asn 1605	Gly	Thr	Thr

Thr Ala His Ala Gly Ser Val Ser Pro Val Val Ser Ala Pro Val 1610 1620

	Leu	Asn 1625	Thr	Leu	Ser	Gln	Gln 1630	Thr	Gly	Gly	Asp	Ser 1635	Leu	Thr	Gln
5	Thr	Ala 1640	Leu	Gln	Gln	Tyr	Glu 1645	Pro	Val	Val	Val	Gly 1650	Ser	Pro	Gln
.10	Trp	His 1655	Asp ·	Glu	Leu	Ala	Gly 1660	Ala	Leu	Lys :		Ile 1665	Ala	Gly	Gly
15	Ser	Pro 1670	Leu	Thr	Gly	Gln	Thr 1675	Gly	Ile	Ser	Asp	Asp 1680	Trp	Pro	Leu
20	Pro	Ser 1685	Gly	Asn	Asn	Gly	Tyr 1690	Leu	Val	Pro		Thr 1695	Asp.	Pro	Asp
	Ser	Pro 1700	Tyr	Leu	Ile	Thr	Val 1705	Asn	Pro	Lys	Leu	Asp 1710	Gly	Leu	Gly
25	Gln	Val 1715.		Ser	His	Leu	Phe 1720	Ala	Gly	Leu	Tyr	Glu 1725	Leu	Leu	Gly
30	Ala	Lys 1730	Pro	Gly	Gln	Ala	Pro 1735	Arg	Ğlu	Thr	Ala	Pro 1740	Ser	Tyr	Thr
35	Asp	Glu 1745	Lys	Gln	Phe	Leu	Gly 1750	Ser	Ser	Tyr	Phe	Leu 1755	Asp	Arg	Leu
40	Gly	Leu 1760	Lys	Pro	Glu	Lys ·	Asp 1765	Tyr	Arg	Phe	Leu	Gly 1770		Ala	Val
	Phe	Asp 1775	Thr	Arg	Tyr	Val	Ser 1780	Asn ·	Ala	Val	Leu	Ser 1785	Arg	Thr	Gly
45	Ser	Arg 1790	Tyr	Leu	Asn	Gly	Leu 1795	Gly	Ser	Asp	Thr	Glu 1800	Gln	Met	Arg
50	Tyr	Leu 1805	Met	Asp	Asn	Ala	Ala 1810	Arg	Gln	Gln	Lys	Gly 1815	Leu	Gly.	Leu

5	Glu	Phe 1820	Gly	Val	Ala	Leu	Thr 1825	Ala	Glų	Gln	Ile	Ala 1830	Gln	Leu	Asp
	Gly	Ser 1835			Trp		Glu 1840	Ser	Val	Thr	Ile	Asn 1845	Gly	Gln	Thr
10	Val	Met 1850	Val	Pro	Lys	Leu	Tyr 1855	Leu	Ser	Pro	Glu	Asp 1860	Ile	Thr	Leu
15	His	Asn 1865	Gly	Ser	Val	Ile	Ser 1870	gly	Asn	Asn	Val	Gln 1875	Leu	Ala	Gly
20	Gly	Asn 1880	Ile	Thr	Asn	Ser	Gly 1885	Gly	Ser	Ile	Asn	Ala 1890	Gln	Asn	Asp ·
25	Leu	Ser 1895	Leu	Asp	Ser	Ser	Gly 1900	Tyr	Ile	Asp	Asn	Leu 1905	Asn	Ala	Gly
	Leu	Ile 1910	Ser	Ala	Gly	Gly	Ser 1915	Leu	Asp	Leu	Ser	Ala 1920	Ile	Gly	Asp
30	Ile	Ser 1925	Asn	Ile	Ser	Ser	Val 1930	Ile	Ser	Gly	Lys	Thr 1935	Val	Gln	Leu
35	Glu	Ser 1940	Val	Ser	Gly	Asn	Ile 1945	Ser	Asn	Ile	Thr	Arg 1950	Arg	Gln	Gln
40	Trp	Asn 1955	Ala	Gly	Ser	Asp	Ser 1960	Gln	Tyr	Gly	Gly	Val 1965	His	Leu	Ser
45	Gly	Thr 1970	Asp	Thr	Gly	Pro	Val 1975	Ala	Thr	Ile	Lys	Gly 1980	Thr	Asp	Ser
	Leu	Ser 1985	Leu	Asp	Ala	Gly	Lys 1990	Asn	Ile	Asp	ile	Thr 1995	Gly	Ala	Thr
50	Val	Ser	Ser	Gly	Gly	Asp	Leu	Gly	Met	Ser	Ala	Gly	Asn	Asp	Ile

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		2000					2005					2010			
5	Asn	Ile 2015	Ala	Ala	Asn	Leu	Ile 2020	Ser	Gly	Ser	Lys	Ser 2025	Gln	Ser	Gly
10	Phe	Trp 2030	His	Thr	Asp	Asp	Asn 2035	Ser	Ser	Ser	Ser	Thr 2040	Thr	Ser	Gln
	Gly ·	Ser 2045	Ser	Ile	Ser	Ala	Gly 2050	Gly	Asn	Leu	Ala	Met 2055	Ala	Ala	Gly
15	His	Asn 2060	Leu	Asp	Val	Thr	Ala 2065	Ser	Ser	Val	Ser	A l a 2070	Gly	His	Ser
20	Ala	Leu 2075	Leu	Ser	Cys	Arg	Ser 2080	Arg	Pro	Ser	Leu	G l u - 2O85	Cys	Ser	Gln
2:5	Gly	Lys 2090	Ala	Lys	Thr	Ser	Arg 2095	Asn	Gly	Arg	Ser	Glu 2100	Ser	His	Glu
30	Ser	His 2105	Ala	Ala	Val	Ser	Thr 2110	Val	Thr	Ala	Gly	Asp 2115	Asn	Phe	Leu
	Leu	Val 2120	Ala	Gly	Arg	Asp	Ile 2125	Ala	Ser	Gln	Ala	Ala 2130	Gly	Met	Ala
35	Ala	Glu 2135	Asn	Asn	Val	Vaļ	Ile 2140	Arg	Gly	Gly	Arg	Asp 2145	Val	Asn	Leu
40	Val	Ala 2150	Glu	Ser	Ala	Gly	Ala 2155	Gly	Asp	Ser	Tyr	Thr 21.60	Ser	Lys	Lys
45	Lys	Lys 2165	Glu	Ile	Asn	Glu	Thr 2170	Val	Arg	Gln	Gln	GLy 2175	Thr	Glu	Ile
50	Ala	Ser 2180	Gly	Gly	Asp	Thr	Thr 2185	Val	Asn	Ala	Gly	Arg 2190	Asp	Ile	Thr

	Ala	Val 2195	Ala	Ser	Ser	Val	Thr 2200	Ala	Thr	Gly	Asn	Ile 2205	Ser	Val	Asn
5	Ala	Gly 2210	Arg	Asp	Val	Ala	Leu 2215	Thr	Thr	Ala	Thr	Glu 2220	Ser	Asp	Туr
10	His	Tyr 2225	Leu	Glu	Thr	Lys	Lys 2230	Lys	Ser	Gly	Gly	Phe 2235	Leu	Ser	Lys
15	Lys	Thr 2240	Thr	Arg	Thr	Ile	Ser 2245	Glu :	Asp	Ser	Ala	Thr 2250	Arg	Glu	Ala
	Gly	Ser 2255	Leu	Leu	\$er	Gly	Asn 2260	Arg	Val	Thr	Val	Asn 2265	Ala	Gly	Asp
20	Asn	Leu 2270	Thr	Val	Glu	Gly	Ser 2275	Asp	Val	Val	Ala	Asp 2280	Arg	Asp	Val
25	Ser	Leu 2285	Ala	Ala	Gly	Asn	His 2290	Val	Asp	Val	Leu	Ala 2295	Aļa	Thr	Ser
30	Thr	Asp 2300	Thr	Ser	Trp	Arg	Phe 2305	Lys	Glu	Thr	Lys	Lys 2310	Ser	Gly	L eu
35	Met	Gly 2315	Thr	Gly	Gly	Ile	Gly 2320	Phe	Thr	Ile	Gly	Ser 2325	Ser	Lys	T hr
	Thr	His 2330	Asp	Arg	Arg	Glu	Ala 2335	Gly	Thr	Thr	Gln	Ser 2340	Gln	Ser	Ala
40	Ser	Thr 2345	Ile	Gly	Ser	Thr	Ala 2350	Gly	Asn	Val	Ser	Ile 2355	Thr	Ala	Gly
45	Lys	Gln 2360	Ala	His	Ile	Ser	Gly 2365	Ser	Asp	Val	Ile	Ala 2370	Asn	Arg	Asp
50	Ile	Ser 2375	Ile	Thr	Gly	Asp	Ser 2380	Val	Val	Val	Asp	Pro 2385	Gly [.]	His	Asp

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	Arg	Arg 2390	Thr	Val	Asp	Glu	Lys 2395	Phe	Glu	Gln	Lys	Lys 2400	Ser	Gly	Leu
	Thr	Val 2405	Ala	Leu	Ser	Gly	Thr 2410	Val	Gly	Ser	Ala	Ile 2415	Asn	Asn	Ala
10	Val	Thr 2420	Ser	Ala	Gln	Glu	Thr 2425	Lys	Glu	Ser	Ser	Asp 2430	Ser	Arg	Leu
15	_	Ala 2435	Leu	Gln	Ala	Thr	Lys 2440	Thr	Ala	Leu	Ser	Gly 2445	Val	Gln	Ala
20	Gly	Gln 2450	Ala	Ala	Thr	Met	Ala 2455	Ser	Ala	Thr	Gly	Asp 2460	Pro	Asn	Ala
	Gly	Val 2465	Ser	Leu	Ser	Leu	Thr 2470	Thr	Gln	Lys	Ser	Lys 2475	Ser	Gln	Gln
25	Ḥis	Ser 2480	Glu	Ser	Asp	Thr	Val 2485	Ser	Gly	Ser	Thr	Leu 2490	Asn	Ala	Gly
30	Asn	Asn 2495	Leu	Ser	Val	Val	Ala 2500	Thr	Gly	Lys	Asn	Arg 2505	Gly	Asp	Asn
35	Arg	Gly 2510	Asp	Ile	Val,	Ile	Ala 2515	Gly	Ser	Gln	Leu	Lys 2520	Ala	Gly	Gly
40	Asn	Thr 2525	Ser	Leu	Asp	Ala	Ala 2530	Asn	Asp	Ile	Leu	Leu 2535	Ser	Gly	Ala
	Ala	Asn 2540	Thr	Gln	Lys	Thr	Thr 2545	Gly	Arg	Asn	Ser	Ser 2550	Ser	Gly	Gly
45	Gly	Val 2555	Gly	Val	Ser	Ile	Gly 2560	Ala	Gly	Lys	Gly	Ala 2565	Gly	Ile	Ser
50	Ala	Phe 2570	Ala	Ser	Val	Asn	Ala 2575	Ala	Lys	Gly	Arg	Glu 2580	Lys	Gly	Asn

5	Gly	Thr 2585	Thr	Thr	Asp	Lys	Thr 2590		Thr	Ile	Asn	Ser 2595	Gly	Arg	Asp
	Thr	Val 2600	Leu	Asn	Gly	Ala	Gln 2605	Val	Asn	Gly	Asn	Arg 2610	Ile	Ile	Ala
10	Asp	Val 2615	Gly	His	Asp	Leu	Leu 2620	Ile	Ser	Ser	Gln	Gln 2625	Asp	Thr	Ser
15	Lys	Tyr 2630	Asp	Ser	Lys	Gln	Thr 2635	Ser	Val	Ala	Ala	Gly 2640	Gly	Ser	Phe
20	Thr	Phe 2645	Gly	Ser	Met	Thr	Gly 2650	Ser	Gl _, y	Tyr	Ile	Ala 2655	Ala	Ser	Arg
25	Åsp	Lys 2660	Met	Lys	Ser	Arg	Phe 2665	Asp	Ser	Val	Ala	Glu 2670	Gln	Thr	Gly
	Met	Phe 2675		Arg	Val	Met	Val 2680	Ala	Ser	Thr	Ser	Gln 2685	Trp	Val	Asn
30	Ile	Pro 2690	Asn	Trp	Met	Val	Arg 2695	Ser	Leu	Pro	His	Cys 2700	His	Thr	Gly
35	Glu	Lys 2705	Pro	Pro	Gly	Tyr	Arg 2710		Leu	Gly	Leu	Val 2715	Thr	Leu	Gln
40	Arg	Ser 2720	. –	Ile	Ile	Lys	Ser 2725		His	Arg	Trp	Asn 2730	Gln	Ser	
45	<210 44)> 44	4 <21	L1>	321	<212	2> PI	RT <2	213>	Esc	cheri	ichia	coli	_ <4	100>
.5	Met Asn 1	Met I		-	iys T	Thr]	(le Ph	ne II	Le Le		nr L∈	eu Phe	e Sei	Gl ₃	7

	V	VO 200	5/09782	23				152/3	70				PCT	EP200	5/00391	72
	Val	Ile	Ala	Ala	Thr	Val	Glu			Phe	Glu	Asn	Glu	Gln	Tyr	
	Asn			20					25			÷		30		
5																
3	-	Ala	Tyr	Arg	Ser	Ala	Asp	Val	Phe	Met	Pro	Tyr	Ile	Lys	Ser	
	Asn	•	35		*			40					45			
10												•				
	Phe Met	Asn	Pro	Val	Thr	Asp	Ser	Ala	Leu	Asn	Val	Ser	Leu	Thr	Tyr	
		50	,		•		55					60				
15	П	Gln	7\	Cln	T-172	C1 x 2	T ***C	Tvc	ដា។ ១	T 175	Luc	Thr	Ser	Glu	7) s n	
	Arg	GTII	Asp	GTII	тйт	_	ту	пуз		пуз		7117	per	Giu	Asp	0.0
	65				•	70					75				•	80
20	Phe	Lys	Thr	Asn	Arg	Asp	Arg	Ile	Glu	Leu	Tyr	Leu	Lys	Gly	Tyr	•
	Thr				85					90		•			95	
25																
23		Asn	Arg	Gly	Ala	Tyr	Ser	Phe	Ser	Pro	Ser	Ala	Gly	Phe	Arg	
	Tyr			100				٠	105					110		
30					•											
	Glu Trp	Ser	Trp	Asp	Val	Asn	Tyr	Asp	Asn	Pro	Lys	Lys	Gln	Asp	Lys	
			115					120					125			
35	Lvs	Leu	Glu	T.eu	Ara	Phe	Ψvr	Pro	Asn	Met	Thr	Tvr	Tivs	Leu	Asn	
	Asp		OIU	Lea	1119	1110	135		11011	1100		140	-1-			
4.0		130					133					140				
40	Gln	Leu	Ser	Leu	Tyr	Met	Asn	Gly	Phe	Val	Ala	Pro	Val	Phe	Phe	
	Lys 145					150					155					
45	160														•	
														'		

Thr Gln Glu Ser Arg Lys Asp Asn Asn Tyr Val Lys Gly Lys
Leu
165 170 175

15	3/37

*	_	Ala	Lys	Arg	Tyr	Asn	Asn	Asp	Tyr	Tyr	Gln	Glu	Leu	Gln	Ile
•	Leu			180					185					190	
5	· Gl v	Val	Ara	Tvr	Lys	Phe	Asn	Asn	Asp	Asn	Thr	Leu	Trp	Ala	Ser
	Val		195	-1-	2			200	1				205		
10															
	Tyr Trp	Asn	Glu	Arg	Lys	Tyr		Gln	His	Ser	Ser		Tyr	Asp	Arg
		210					215					220			
15		Leu	Arg	Gly	Gly	Tyr	Asp	Phe	Lys	Val	Thr	Glu	Glu	Phe	Val
	Leu 225					230					235				
20	240														
•	Ser Glu	Pro	Phe	Ile	Arg	Tyr	Asp	Leu	Ser	Tyr	Arg	Glu	Lys	Asn	Leu
25	GLU				245					250.					255
20	Ser	Thr	Ser	Asn	Asn	Glv	Leu	Ser	Lys	Asn	Asn	Lys	Glu	Ile	Arq
*	Thr			260					265			_		270	,
30										ž.			•		
	Gly Gly	Ala	Ser	Phe	Ser	Tyr	Lys	Ile	Ile	Pro	Ser	Val	Lys	Leu	Val
35 .			275					280					285		
		Ile	Tyr	Arg	Gln	Thr	Thr	Asn	Ile	Glu	Asn	Tyr	Tyr	Gly	Glu
40	His	290					295				-	300			
40	C a sa	Cl.,	7)	Tira	7.00	7) 25 05	Mot	Dho	m	Tuc	T 011	Clu	Tlo	7\	Tuc
	Thr	GIU	Asp	ήλε	Asn	310	Mec	rne	тут	пуз	315	Grð	116	ASII	пуз
45	320					010			•						
	Phe												2		

WO 2005/097823 PCT/EP2005/003972 154/370

<210>	45 <211>	587 <212>	PRT <213>	Escherichia coli	<400>
45					

Met Gln His Arg Gln Lys Asn Ile Leu Thr Lys Thr Ser Leu Leu 5 Ser 1 5 10 15

Arg Ala Leu Ser Val Pro Cys Cys Asp Met Phe Arg Arg Gly Ser 10 Pro 20 25 30

Trp Ile Cys Tyr Leu Ser Leu Ser Val Phe Ser Gly Cys Phe Ile 15 Pro 35 40 45

Ala Phe Ser Ser Pro Ala Ala Met Leu Ser Pro Gly Asp Arg Ser 20 Ala 50 55 60

Ile Gln Gln Gln Gln Gln Leu Leu Asp Glu Asn Gln Arg Gln

Arg
65 70 75 80

Asp Ala Leu Glu Arg Pro Leu Thr Ile Thr Pro Ser Pro Glu Thr 30 Ser 85 90 95

Gly Ala Thr Arg Leu Thr Ser Ala Glu Thr Asp Arg Leu Val Pro
40 Trp
115 120 125

Val Asn Gln Cys Leu Asn Ile Thr Gly Leu Thr Ala Val Thr Asp 45 Ala 130 135 140

Val Thr Asp Gly Tyr Ile Arg Arg Gly Tyr Ile Thr Ser Arg Ala $50\,$ Phe

14 5 150 155 16 O

5 Leu Thr Glu Gln Asp Leu Ser Gly Gly Val Leu His Ile Thr Val Met

165 170 175

10 Glu Gly Arg Leu Gln Gln Ile Arg Ala Glu Gly Ala Asp Leu Pro Ala 180 185 190

15 Arg Thr Leu Lys Met Val Phe Pro Gly Met Glu Gly Lys Val Leu Asn 200 205

20 Leu Arg Asp Ile Glu Gln Gly Met Glu Gln Ile Asn Arg Leu Arg
Thr
210 215 220

25 Glu Pro Val Gln Ile Glu Ile Ser Pro Gly Asp Arg Glu Gly Trp Ser 225 230 235 240

30
Val Val Thr Leu Thr Ala Leu Pro Glu Trp Pro Val Thr Gly Ser Val

245
250
255

Gly Ile Asp Asn Ser Gly Gln Lys Ser Thr Gly Thr Gly Gln Leu Asn
260 265 270

Gly Val Leu Ser Phe Asn Asn Pro Leu Gly Leu Ala Asp Asn Trp Phe
275 280 285

Val Ser Gly Gly Arg Ser Ser Asp Phe Ser Val Ser His Asp Ala Arg
290 295 300

	W	/ O 2 00:	5/09782	23				156/3	70		,		PCT/	EP2005	5/003972
	Asn Asp	Phe	Ala	Ala	Gly	,Val	Ser	Leu	Pro	Tyr	Gly	Tyr	Thr	Leu	Val
5	305 320					31 0					315				
	Tyr Gly	Thr	Tyr	Ser	Trp	Ser	Asp	Tyr	Leu	Ser	Thr	Ile	Asp	Asn	Arg
10	Gry				325					330					335
	Trp Leu	Arg	Trp	Arg	Ser	Thr	Gly	Asp	·Leu	Gln	Thr	His	Arg	Leu	Gļy
15	Ecu			340					345					350	
	Ser Gly	His	Val	Leu	Phe	Arg	Asn	Gly	Asp	Met	Lys	Thr	Ala	Leu	Thr
20			355	G.				360				*	365		
	Gly Leu	Leu	Gln	His	Arg	Ile	Ile	His	Asn	Tyr	Leu	Asp	Asp	Val	Leu
25		370		,			375			-		380			
	Gln His	Gly	Ser	Ser	Arg	Lys	Leu	Thṛ	Ser	Phe	Ser	Val	Gly	Leu	Asn .
30	385 400				i (i)	390					395				

Thr His Lys Phe Leu Gly Gly Val Gly Thr Leu Asn Pro Val Phe

Arg Gly Met Pro Trp Phe Gly Ala Glu Ser Asp His Gly Lys Arg

Asp Leu Pro Val Asn Glm Phe Arg Lys Trp Ser Val Ser Ala Ser

Gln Arg Pro Val Thr Asp Arg Val Trp Trp Leu Thr Ser Ala Tyr

Thr

Gly

Phe

Ala

110 2000/05 1020		
	157/370	_

* 5	Gln Gly 465 480	Trp	Ser	Pro	Asp	Arg 470	Leu	His	Gl y	Val	Glu 475	Gln	Leu	Ser	Leu
10	Gly Asn	Glu	Ser	Ser	Val 485	Arg	Gly	Phe	Lys	Asp 490	Gln	Tyr	Ile	Ser	Gly 495
15	Asn Leu	Gly	Gly	Tyr 500	Leu	Arg	Asn	Glu	Lę u 50 5	Ser	Trp	Ser	Leu	Phe 510	Ser .
20	Pro Trp	Tyr	Val	Gly	Thr	Val	Arg	Ala 520	Val	Ala	Ala	Leu	Asp 525	Gly	Gly
25	Leu Gly	His 530	Ser	Asp	Ser	Asp	Asp 535	Pro	Tyr	Ser	Ser	Gly 540	Thr	Leu	Trp
30	Ala Phe 545 560	Ala	Ala	Gly	Leu	Ser 550	Thr	Thr	Ser	Gly	His 555	Val	Ser	Gly	Ser
35	Thr His	Ala	Gly	Leu	Pro 565	Leu	Val	Tyr	Pro	Asp 570	Trp	Leu	Ala	Pro	Asp 575
40	Leu	Thr		Tyr 580	Trp	Arg	Val	Ala	Val 585	Ala	Phe	٠			
45	<210 46)> 4	16 <2	211>	744	<21	.2>	PRT	<213	3> E	Esche	erich	ia c	oli	<400>
+2	Met Ile 1	Asn	Lys	His	Thr _. 5	Leu	Leu	Leu	Thx		Leu			Asn	Leu
50															

	V	V O 2 00	5/0978	323				150	/2 7 0				PCT	EP200	5/00391	72
			*					158/	3/0							
	Cys Gln	Thr	Pro	Val	Phe	Ala	Gln	Asn	Trp	Gln	Val	Ala	Thr	Phe	Gly	
	0111			20					25					30		
5		•								4				,		
	Ser Gly	Thr	Asp	Leu	Asn	Phe	Ser	Ser	Leu	Ile	Asp	Ser	Ala	Lys	Ile	
	_		35					40					45			
10																
,	Arg Gly	Asn	Asn	Ala	Trp	Leu	Ala	Gly	Asn	Asn	Asn	₽he	Leu	Glu	Ala	
	1	50			*		55					60				
15		_,		1	_	_	 1	. .	7 0.1	7.1	~~ T		G	70	G 3	
•	Lys Gly	Phe	Tyr	Thr	Leu	Pro	Thr	Asp	Phe	Phe	TTE	Glu	Ser	Arg	GTA	
	65					70					75			•		8 (
20 .	ų.							ı								
	Lys Val	Ile	Ala	Asn	Ser	His	Asp	Gly	Met	Thr	Val	Phe	Tyr	Thr	Ile	
					85 .					90					95	
25																
	Pro Gln	Val	Thr	Gln	Thr	Phe	Arg	Leu	Glu	Ala	Asp	Lęu	Thr	Leu	Glu	
	OIII			100		•			105					110		
30																
		Gly	Pro	Glu	Val	Asn	Gly	Lys	Ser	Pro	Ala	Gly	Gln	Glu	Gly	
	Ala		115					120					125			
35																
33		Leu	Phe	Val	Arg	Asp	Iļle	Ile	Gly	Pro	Gln	Arg	Gln	Glu	Pro	
	Gln	130					135					140	-			
40			•													
40		Ala	Gly	Thr	Glu	Glu	Tyr	Pro	Gln	Ala	Ser	Ælsn	Ile	Leu	Met	
	Asn 145					150					155					
45	160															
	7\] ~	Dho	Tla	ሞኮ∽	Gl n	Aen	T.ve	T.ve	Zen	Asn	Asn	Leu	Val	Gln	Tle	
	Thr	LIIG	TTE	7.117			пуз	пуз	ASII		11011	<u></u> cu	vul	- - - 11		
					1.65					170					175	

	Ser Ile		Val	Arg	Glu	Gly	Val	Ile	Lys	Thr	Trp	Gly	Asn	Glu	Gly
_				180			,		185					190	
5			Lys	Lys	Gln	Pro	Ile	Ile	Glu	Asn	Ile	Asn	Phe	Thr	Gln
	Lys		195					200	•				205		
10	Arg	Asn	Ile	His	Met	Thr	Ile	Glu	Arg	Leu	Pro	Glu [,]	Lvs	Phe	Ile
	Leu						. 215					220	-		
15	ጥኮሎ	7\ 1 ->	Dho	7. a.s.	Th ν	· 7\ a.s.	71 20 00	Ta	C1	7)	<i>C</i> 1	G	5 0	C.I	7.
	Ser 225	EATA	rne	Asp	T11T	230	ALG	тух	GIU	Asn	235	.ser	rrp	Gln	Phe
20	240														•
		Tyr	Ser	Gly	Phe	Met	Äsn	Gln	Leu	Asp	Asn	Asn	Ser	Leu	Ala
25	Ile				245	_				250				•	255
	Gly	Phe	Phe	Ala	Ala	Arg	Asn	Ala	Lys	Leu	Árg	Val	Lys	Asn	Ala
20	Ser	•		260					265					270	
30	Phe	Tive	Pro	Gly	Tage	Pro	T. 211	Val	7\ s.n	Tur	Tvc	Cln	Tou	Thr	C 0 **
	Arg	гуу	275	OTY	пуъ	FLO	пец	280	Yab		пуѕ	GT'II	285	THE	ser
35			•										200	*	
	Gln Ser		Ser	Arg	Val	Arg		Lys	Ala	Pro	Ģlu		Phe	Leu	Ala
40	•	290					295	•				300			
	Pro Asn	Gln	Ser	Val.	Val	Arg	Asn	Ser	Thr	Thr	Leu	Gln	Phe	Leu	Ala
45	305 320	-				310					315				
		,		_ =											
50	Gln Val	Ala	GLY	ITe		Ser	Ile	Asp	Asn		Lys	Gln	Thr	Lys	•
50					325					330					335

Gln Ala Gly Glu Leu Val Gln Phe Pro Val Thr Leu Gln Lys Lys His Asn Asp Phe Thr Val Asn Phe Asn Val Asp Gly Asn Ile Ser Lys Lys Ala Ile Arg Ile Glu Gln Val Lys Ser Asn Leu Thr Asp Pro Tyr Glu Ile Tyr Val Cys Ser Asp Cys Arg Gln Gly Ala Arg Gly Ser Lys Asn 390 -Asp Pro Val Asp Leu Gln Thr Ala Val Lys Phe Val Ala Pro Gly Gly Asn Ile Tyr Leu Asn Asp Gly Gln Tyr His Gly Ile Thr Leu Asp Arg .425 Glu Leu Ser Gly Ile Pro Gly Lys Tyr Lys Thr Ile Ser Ala Ile Asn Pro His Lys Ala Ile Phe Ile Asn Lys Thr Phe Asn Leu Asp Ala Ser Tyr Trp His Leu Lys Ser Val Val Phe Asp Gly Asn Val Asp Asn Gly Asn Asn Lys Pro Ala Tyr Leu Arg Ile Ala Gly Ser Tyr Asn Ile Ile

	Glu Ser	His	Val	Ile	Ala	Arg	Asn	Asn	Asp	Asp	Thr	Gly	Ile	Ser	Ile
5				500					505		•			510	
	Ala	Lys	Asp	Lys	Asn	Arg	Phe	Phe	Trp	Pro	Ala	His	Asn	Leu	Val
10	Leu		515					520					525		
				•											
1.7	Asn Asp		Asp	Ser	Tyr	Asn		Leu	Asp	Leu	Ser		Ile	Asn	Ala
15		530					535					540			
		Phe	Ala	Ala	Lys	Leu	Gly	Val	Gly	Pro	Gly	Asn	Ile	Phe	Arg
20	Gly 545 560					550					555			-	
	300														
25	Cys Lys	İle	Ala	His	Asn	Asn	Ala	Asp	Asp	Gly	Trp	Asp	Leu	Phe	Asn
	4				565					570		,			575
		Glu	Asp	Gly	Pro	Asn	Ala	Ser	Val	Thr	Ile	Glu	Asn	Ser	Val
30	Ala	•	.'	580					585			•		590	
			_			_	·		_		-				
35	Tyr Ser	GLU		Gly	Leu	Pro	Tyr		Lys	Ala	Asp	Ile		Lys	Gly
			595					600					605		
40	Ile Asn	Gly	Asn	Gly	Gly	Glu	Gly	Gln	Pro	Ser	Lys	Ser.	Gln	Val	Ile
.0	11011	610	,				615					620			,
	Ser	Ile	Ala	Ile	Asn.	Asn	Asn	Met	Asp	Glv	Phe	Thr	`Asp	Asn	Phe
45	Asn 625				•	630			_		635		<u>r</u>		
	640								•						
50	Thr	Gly	Ser	Leu	Ile	Val	Arg	Asn	Asn	Ile	Ala	Met	Asn	Asn	Ala
	7\ >0.00														

Arg

	W	O 2005	5/097 82 3	3				162/3	70				PCT/I	EP2005	5/003972
					645					650					655
5	Tyr Ile	Asn	Tyr	Ile 660	Leu	Arg	Thr	Asn	Pro 665	Tyr	Lys	Phe	Pro	Ser 670	Ser
10	Leu Ile	Phe	Asp 675	Asn	Asn	Tyr	Ser	Ile 680	Arg	Asp	Asp	Trp	Glu 685	Asn	Lys
15	Lys Leu	Asp	Phe	Leu	Gly	Asp	Thr 695	Val	Asn	Ser	Val	Asn 700	Tyr	Lys	Leu
20	Val Arg 705 720	Ser	His	Glu	Thr	Gly 710	Pro	Val	Gl'n	Lys	Asp 715	Leu	Phe	Phe	Thr
25	Asp Ile	Asp	Ser	Gly	Asn 725	Ile	Ile	Tyr	Pro	Asp 730	Phe	Phe	Leu	Asn	Ile 735
30	Asn	Lys	Phe	Asn 740	Glx	Thr	Met	Pro						,	
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45	Ser Lys	Val	Phe	Ala 20	Thr	Ala	Lys	Gln	Val	Lys	Leu	Pro	Asn	Asn 30	Ile
	Tyr Met	Val	Asn	Thr	Thr	Glu	Ala	Phe	Ser	Cys	Thr	Glu	Ile	Asp	Gly

Asn Cys Gln Thr Lys Asn Pro Phe Asn Tyr Lys Asp Asn Ser Tyr Val

Phe Val Leu Glu Arg Gly Gly Ala Trp Cys Tyr Asp Tyr Thr Val Ser

Val Leu Asn Leu Lys Thr Gly Lys Ala Gln Met Leu Glu Tyr Lys Asp

Asn Gln Leu Cys Ser Gly Ser Asn Lys Pro Phe Phe Glu Ile Lys

Gly Val Pro Thr Val Gly Val Ile Asp Thr Ser Gly Lys Pro Val Val

Val Ala Leu Asp Lys Leu Lys Thr

<210> 48 <211> 225 <212> PRT <213> Escherichia coli <400>

Met Gln Leu Pro Val Lys Leu Leu Met Ser Leu Ile Ser Leu Val Ser

Val Ile Ala Arg Ala Gly Lys Tyr Lys Asn Tyr Ile Arg Asp Glu Ile

Lys Tyr Trp Arg Tyr Thr Ser Tyr Lys Gly Glu Phe Pro Glu Gly

Phe Thr Asp Glu Lys Phe Ser Ser Ala Ile Tyr Asn Gly Arg Ile Phe

5	Thr Phe 65	Met ·	Lys	Arg	Leu	His	Thr	Leu	Met	Leu	Phe	Leu	Ala	Val	Leu	80
	Thr Cys	Gly	Phe	Asn	Val	Glu	Ala	Ala	Ser	Val	Lys	Gln	Ala	Leu	Ser	
10							•									
	Asp Tyr	Pro	Asn	Ala	Arg	Ala	Glu	Gln	Pro	Gly	Ala	Cys	Pro	Thr	Thr	
15	4			100					105					110		
13	01	T		Q1	G 3	71	73 7	73. 73		_			_	_		
	Ala	ьeu		GIU	Gly	Asp	Ala		Tyr	туѕ	Ala	Ala		Asp	Lys	
20			115				•	120					125			
	Leu	Lys	Pro	Val	Gly	Leu	Ser	Gly	Met	Phe	Glv	Lvs	Glv	Glv	Tvr	
	Met	130		,			135	2				140	- · 1	1	-1-	
25		100					100					T40				
		Gly	Pro	Gly	Gly	Asn	Val	Thr	Pro	Val	Thr	Ile [.]	Asn	Gly	Thr	
	Val 145					150					155					
30	160															
	Trp	Leu	Gln	Gly	Asp	Gly	Cys	Lys	Ala	Asn	Thr	Cys	Glv	Trp	Asp	
35	Phe			_	165	_	-	-		170		•	-		175	
					,											
		Val	Thr	Leu	Tyr	Asn	Pro	Lys	Thr	His	Glu	Val	Val	Gly	Tyr	
40	Arg			180					185					190		
					•			•					•			
	Tyr Ile	Phe	Gly	Leu	Asp	Asp	Pro	Ala	Tyr	Leu	Val.	Trp	Phe	Gly	Glu	
45			195					200					205			ŧ
		77-7	ui-	C1	Dh -	7\ 7 —	M	Т с		T	7\	m	*7-7	71.7 -	71 T -	
~ 0	Val			GLU	Phe	Ата		ьeu	val	туѕ	ASN		vaı	Α⊥а	Ата	
50	•	210					215					220				

Asn 225

5 <210> 49 <211> 721 <212> PRT <213> Escherichia coli <400> 49

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Phe Ile Pro Leu His Ala His Ala Ser Ser Thr Ser Glu Asp Glu
15 Met
20 25 30

Ile Val Thr Gly Asn Thr Ala Ala Asp Thr Thr Asp Ser Ala Ala 20 Gly 35 40 45

Ala Gly Phe Lys Thr Asn Asp Ile Asp Val Gly Pro Leu Gly Thr
Lys
50
55
60

Ser Trp Ile Glu Thr Pro Tyr Ser Ser Thr Thr Val Thr Lys Glu

Met
65 70 75 80

Ile Glu Asn Gln Gln Ala Gln Ser Val Ser Glu Met Leu Lys Tyr 35 Ser 85 90 95

Pro Ser Thr Gln Met Gln Ala Arg Gly Gly Met Asp Val Gly Arg
40 Pro
100 105 110

Gln Ser Arg Gly Met Gln Gly Ser Val Val Ala Asn Ser Arg Leu 45 Asp 115 120 125

Gly Leu Asn Ile Val Ser Thr Thr Ala Phe Pro Val Glu Met Leu 50 Glu 130 135 140

5	Arg Ala 145 160	Met	Asp	Val	Leu	Asn 150	Ser	Leu ·	Thr	Gly	Ala 155	Leu	Tyr	Gly	Pro
10	Ser Glu	Pro	Ala	Gly	Gln 165	Phe	Asn	Phe	Val	Ala 170	Lys	Arg	Pro	Thr	Glu 175
15	Thr Thr	Leu	Arg	Lys 180	Val	Thr :	Leu	Gly	Tyr 185		Ser	Arg	Ser	Ala 190	Phe
20	Gly Gly	His	Ala 195	Asp	Leu	Gly	Gly	His 200	Phe	Asp	Glu	Asn	Lys 205	Arg	Phe
25	Tyr Asp	Arg 210	Val	Asn	Leu	Leu	Asp 215	Gln	Glu	Gly	Glu	Gly 220	Asn	Val	Asp
30	Ser Ile 225 240	Thr	Leu	Arg	Arg	Lys 230	Leu	Val	Ser	Val	Ala 235	Leu	Asp	Trp	Asn
35	Gln Ile	Pro	Gly	Thr	Gln 245	Leu	Gln	Leu	Asp	Ala 250	Ser	His	Tyr	Glu	Phe 255
40	Gln Leu	Lys	Gly	Tyr 260	Val	Gly	Ser	Phe	Asn 265	Tyr	Gly	Pro	Asn	Val 270	Lys
45	Pro Ala	Ser	Ala 275	Pro	Asn	Pro	Lys	Asp 280	Lys	Asn	Leu	Ala	Leu 285	Ser	Thr
50	Gly Tyr	Asn	Asp	Leu	Thr	Ţhr	Asp	Thr	Ile	Ser	Thr	Arg	Leu	Ile	His

290		205	300
290	-	295	300

5	Phe Ala 305 320	Asn	Asp	Asp	Trp	Ser	Met	Asn	Ala	Gly	Val	Gly	Trp	Gln	Gln
10	Asp Gly	Arg	Ala	Met	Arg	Ser	Val	Ser	Ser	Lys	Ile	Leu	Asn	Asn	Gln 335
15	Asp Arg	Ile	Ser	Arg	Ser	Met	Lys	Asp	Ser 345	Thr	Ala	Ala	Gly	Arg 350	Phe
20	Val Ser	Leu	Ser 355	Asn	Th±	Ala	Gly	Leu 360	Asn	Gly	His	Ile	Asp 365	Thr	Gly
25	Ile Leu	Gly 370	His	Asp	Leu	Ser	Leu 375	Ser	Thr	Thr	Gly	Tyr 380	Val	Trp	Ser
30	Tyr Asn 385 400	Ser	Ala	Lys	Gly	Thr 390	Gly	Ser	Ser	Tyr	Ser 395		Gly	Thr	Thr
35	Met Arg	Tyr	His	Pro	Asp 405	Ala	Ile	Asp	Glu	Gln 410	Gly	Asp	Gly	Lys	Ile 415
40	Thr Val	Gly	Gly	Pro 420	Arg	Tyr	Arg	Ser	Ser 425	Val	Asn	Thr	Gln	Gln 430	Ser
45	Thr Phe	Leu	Gly 435	Asp	Thr	Val	Thr	Phe	Thr	Pro	Gln	Trp	Ser 445	Ala	Met

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	Tyr Gly	Leu	Ser	: Gln	Ser	Trp	Leu	Gln	Thr	Lys	Asn	Tyr	Asp	Lys	His
	. –	450					455					460			•
5	70		 .	-		7						,			
	Ala	GIn	Thr	Asn	. GIn	Val	Asp	Glu	Asn	Gly	Leu	Ser	Pro	Asn	Ala
	465 480					470					475				
10															
4	Leu Ala	Met	Tyr	Lys	Ile	Thr	Pro	Asn	Thr	Met	Ala	Tyr	Vạl	Ser	Tyr
15		•			485				-	490					495
15		~									5 15				
	Asp Lys	Ser	Leu	Glu	Gln	Gly	Gly	Thr	Ala	Pro	Thr	Asp	Glu	Ser	Val
20 ·				500					505					510	
•	Asn	Δla	Glv	Gln	Thr	Lou	7) cn	Dxo	П.т.	. 7) 20 00	C	T	0.7		
	Val	211.0		OTII	TIIT	пеп	Asn		т Хт.	Arg	ser	гуѕ		Tyr	GLu
25	,		515		•			520					525		
	Gly	Leu	Lys	Ser	Asp	Ile	Gly	Glu	Met	Asn	Leu	Glv	Ala	Ala	Leu
	Phe	530					535					540			
30								·							
	Arg	Leu	Glu	Arg	Pro	Phe	Ala	Tyr	Leu	Asp	Thr	Asp	Asn	Val	Tyr
	Lys 545					550					555				-
35	560			Ť											
٠.	G] 11	Gln	Glv	Asn	Gln	[cV	A en	7) G D	Cl w	T 011	Clu	T 011	mb	7.1 _	70 T . •
40	Gly	OIII	C II y	Woii	•	vai	Asn	. AS≢I	GTĀ		GLU	теп	Inr	Ата	Ala
40					565					570					575
	Asn	Val	Trp	Gln	Gly	Leu	Asn	Ile	Tvr	Ser	Glv	Val	Thr	Phe	T.e.i
45	Asp			580	_				585		.				
-														590	
	Pro	Lys	Leu	Lys	Asp	Thr	Ala	Asn	Ala	Ser	Thr	Ser	Asn	Lys	Gln
50	Val		595					600					6Ö5		
				•							,				

	1.00/2
	169/3

Val Gly Val Pro Lys Val Gln Ala Asn Leu Leu Ala Glu Tyr Ser Leu Pro Ser Ile Pro Glu Trp Val Tyr Ser Ala Asn Val His Tyr Thr Gly Lys Arg Ala Ala Asn Asp Thr Asn Thr Ser Tyr Ala Ser Ser Tyr. Thr Thr Trp Asp Leu Gly Thr Arg Tyr Thr Thr Lys Val Ser Asn Val Pro. Thr Thr Phe Arg Val Val Val Asn Asn Val Phe Asp Lys His Tyr Trp Ala Ser Ile Phe Pro Ser Gly Thr Asp Gly Asp Asn Gly Ser Pro Ser Ala Phe Ile Gly Gly Gly Arg Glu Val Arg Ala Ser Val Thr Phe Asp Phe 50 <211> 669 <212> PRT <213> Escherichia coli <400> <210>

Met Lys Asn Ile Thr Leu Trp Gln Arg Leu Arg Gln Val Ser Ile Ser 1.0

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				•				170/	370							
	Thr Ile	Ser	Leu	Arg	Cys	Ala	Phe	Leu	Met	Gly	Ala	Leu	Leu	Thr	Leu	
				20					25			v		30		
5	Val	Ser	Ser	Val	Ser	Leu	Týr	Ser	Trp	His	Glu	Gln	Ser	Ser	Gln	
ı	Ile		35					40					45			
								0					40			
10	Arg Leu	Tyr	Ser	Leu	Asp	Lys	Tyr	Phe	Pro	Arg	Ile	His	Ser	Ala	Phe	
		50					55					60				
15				-												. •
	Leu	Glu	Gly	Asn	Leu		Leu	Val	Val	Asp		Leu	Asn	Glu	Phe	
	65			e ,		70					75		•			80
20																
,	Gln Ile	Ala	Pro	Asn	Thr	Thr	Val	Arg	Leu	Gln	Leu	Arg	Thr	Gln	Ile	
					85					90					95	
25														•		y.
•	Gln Arg	His	Leu	Asp	Thr	Ile	Glu	Arg	Leu	.Ser	Arg	Gly	Leu	Ser	Ser	
	9			100					105					110	*	
30																
30	Glu Leu	Arg	Gln	Gln	Leu	Thr	Val	Ile	Leu	Gln	Asp	Ser	Arg	Ser	Leu	
			115					120					125			
35			•													
	Ser Lys	Glu	Leu	Asp	Arg	Ala	Ĺeu	Tyr	Asn	Met	Phe	Leu	Leu	Arg	Glu	
	-1-	130					135			-		ユ40				
40														•		
	Val Thr	Ser	Glu	Leu	Ser	Ala	Arg	Ile	Asp	Trp	Leu	His	Asp	Asp	Phe	
	145			•		150					155					
45	160												-			
		_											-			
	Thr Thr	Glu	Leu	Asn	Ser	Leu	Val	Gln	Asp	Phe.	Thr	Trp	Gln	Gln	Gly	
50					165					170					175	
50																

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	Leu Leu	Leu	Asp	Gln 180	Ile	Ala	Ser	Arg	Gln 185	Gly	Asp	Thr	Ala	Gln 190	Tyr
5	Lys	Arg	Ser	Arg	Glu	Val	Gln	Asn		Gln	Gln	Gln	Val		Thr
•	Leu		195					200					205		J. 1. J.
10	Ala Asn	Arg	Ile	Glu	Asn	Gln	Ile	Val	Asp	Asp	Leu	Arg	Asp	Arg	Leu
15	,	210					215					220			
13	Leu	Leu	Lys	Ser	Gly		Asp	Asp	Asp	Ile		Val	Glu	Thr	His
20	225 240					230					235		·		
	Arg Met	Tyr	Phe	Glu	Asn	Leu	Lys	Lys	Thr	Ala	Asp	Glu	Asn	Ile	Arg
25					245					250				•	25 5
	Leu Glu	Asp	Asp	Trp 260	Pro	Gly	Thr	Ile		Leu	Arg	Gln	Thr		Asp
30	.	7	7						265					270	
	Arg	ьeu	275	Met	GTÀ	TTE	Val	280	Asn	ГÀЗ	Met	Pro	Asp 285	Thr	Met
35	C1	П	77 7	71 7	70 T	G]	T		-	0. 3					
40	Arg	290	Val	Ald	ALA	GTU	ьуs 295	АІа	ьеи	GLU	Asp	300	Ser	Arg	Thx
40	Glu	Ala	Thr	Gln	Glv	Ara	Phe	Ara	Thr	Leu	Leu	Glu	Ala	Gln	Leu
45	Gly 305 320		•			310		•			315				
		Thr	His	Gln	Gln	Met	Gln	Met	Phe	Asn	Gln	Arg	Met	Glu	Gln
50	Ile	*			325		•			330					335

Val His Val Ser Gly Gly Leu Ile Leu Val Ala Thr Ala Leu Ala Leu Leu Ala Trp Val Phe Asn His Tyr Phe Ile Arg Ser Arg Leu Val Lys Arg Phe Thr Leu Leu Asn Gln Ala Val Val Gln Ile Gly Leu Gly Gly Thr Glu Thr Thr Ile Pro Val Tyr Gly Asn Asp Glu Leu Gly Arq Ile Ala Gly Leu Leu Arg His Thr Leu Gly Gln Leu Asn Val Gln Lys Gln Gln Leu Glu Gln Glu Ile Thr Asp. Arg Lys Val Ile Glu Ala Asp Leu Arg Ala Thr Gln Asp Glu Leu Ile Gln Thr Ala Lys Leu Ala Val Val Gly Gln Thr Met Thr Leu Ala His Glu Ile Asn Gln Pro Asn Ala Leu Ser Met Tyr Leu Phe Thr Ala Arg Arg Ala Ile Glu Gln Thr Gln Lys Glu Gln Ala Ser Met Met Leu Gly Lys Ala Glu Gly Val

5	Ile Arg	Ser	Arg	Ile 500	Asp	Ala	Ile		Arg	Ser	Leu	Arg	Gln	Phe	Thr
														010	
	Arg Met	Ala	Glu	Leu	Glu	Thr	Ser	Leu	His	Ala	Val	Asp	Leu	Ala	Gln
10			515	•				520				=	525		
*	Phe Gln	Ser	Ala	Ala	Trp	Glu	Leu	Leu	Ala	Met	Arg	His	Arg	Ser	Leu
.15		530					535					540			
3	Ala Glu	Thr	Leu	Val	Leu	Pro	Gln	Gly	Thr	Ala	Thr	Val	Ser	Gly	Asp
20	545 560					550					555				
25	Val Asp	Arg	Thr	Gln	Gln	Val	Leu	Val	Asn	Val	Leu	Ala	Asn	Ala	Leu
23	· ·				565	. 10				570					575
30	Val Gly	Cys	Gly	Gln	Gly	Ala	Val	Ile	Thr	Val	Asn	Trp	Gln	Met	Gln
	-			580					585					590	•
35	Lys Glu-	Thr	Leu	Asn	Val	Phe	Ile	Gly	Asp	Asn	Gly	Pro	Gly	Trp	Pro
	-		595					600					605		
40	Ala Val	Leu	Leu	Pro	Ser	Leu	Leu	Lys	Pro	Phe	Thr	Thr	Ser	Lys	Glu
		610					615					620			
45	Gly Met	Leu	Gly	Ile	Gly	Leu	Ser	Ile	Cys	Val	Ser	Leu	Met	Glu	Gln
	625 640					630					635				
5O	Lys Val	Gly	Glu	Leu	Arg	Leu	Ala	Ser	Thr	Met	Thr	Arg	Asn	Ala	Cys

655

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Val Leu Gln Phe Arg Leu Thr Asp Val Glu Asp Ala Lys 660 665

645

<210> 51 <211> 753 <212> PRT <213> Escherichia coli <400> 51

Met Asn Val Ile Lys Leu Ala Ile Gly Ser Gly Ile Leu Leu Ser

1 5 10 15

Cys Gly Ala Tyr Ser Gln Ser Ile Ser Glu Lys Thr Asn Ser Asp Lys
20
25
30

20 Lys Gly Ala Ala Glu Phe Ser Pro Leu Ser Val Ser Val Gly Lys Thr 35 40 45

25
Thr Ser Glu Gln Glu Ala Leu Glu Lys Thr Gly Ala Thr Ser Ser Arg
50
55
60

30
Thr Thr Asp Lys Asn Leu Gln Ser Leu Asp Ala Thr Val Arg Ser Met
65
70
75
80

Pro Gly Thr Tyr Thr Gln Ile Asp Pro Gly Gln Gly Ala Ile Ser Val

85
90
95

Asn Ile Arg Gly Met Ser Gly Phe Gly Arg Val Asn Thr Met Val Asp

100 105 110

Gly Ile Thr Gln Ser Phe Tyr Gly Thr Ser Thr Ser Gly Thr Thr
Thr
115 120 125

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								1/5/3	70								
	His Leu	Gly	Ser	Thr	Asn	Asn	Met	Ala	Gly	Val	Leu	Ile	Asp	Pro	Asn		
		130					·135					140					
5	Leu Gly 145 160	Val	Ala	Val	Asp	Val 150	Thr	Arg	Gly	Asp	Ser 155	Ser	Gly	Ser	Glu		
10					*(•							
	Ile Asp	Asn	Ala	Leu	Ala	Gly	Ser	Ala	Asn	Met	Arg	Thr	Ile	Gly	Val		
15					165					170					175		
	Asp Ser	Val	Ile	Phe	Asn	Gly	Asn	Thr	Tyr	Gly	Leu	Arg	Ser	Arg	Phe		
20				180					185		-			190			
	Val Gly	Gly	Ser	Asn	Gly	Leu	Gly	Arg	Ser	Gly	Met	Ile	Ala	Ĺeu	Gly		
25	CIJ	•	195					20,0					205				
	Lys Ala	Ser	Asp	Ala	Phe	Thr	Asp	Thr	Gly	Ser	Ile	Gly	Val	Met	Ala	-	
30	AIA	210					215			•		220					
	Val Ile 225	Ser	Gly	Ser	Ser	Val 230	Tyr	Ser	Asn	Phe		Asn	Gly	Ser	Gly		
35	240					230					235						
	Asn Lys	Ser	Lys	Glu	Phe	Gly	Tyr	Asp	Lys	Tyr	Met	Lys	Gln	Asn	Pro		
40	пуз				245					250					255		
	Ser Phe	Gln	Leu	Tyr	Lys	Met	Asp	Ile	Arg	Pro	Asp	Glu	Phe	Asn	Ser		
45	1110			260					265					270			
	Glu Ile	Leu	Ser	Ala	Arg	Thr	Tyr	Glu	Asn	Lys	Phe	Thr	Arg	Arg	Asp		
50	116		275					280					285				

'Thr Ser Asp Asp Tyr Tyr Ile Lys Tyr His Tyr Thr Pro Phe Ser Glu Leu Ile Asp Phe Asn Val Thr Ala Ser Thr Ser Arg Gly Asn Gln Tyr Arg Asp Gly Ser Leu Tyr Thr Phe Tyr Lys Thr Ser Ala Gln Arg Ser Asp Ala Leu Asp Ile Asn Asn Thr Ser Arg Phe Thr Val Ala 2.0 Asp Asn Asp Leu Glu Phe Met Leu Gly Ser Lys Leu Met Arg Thr Tyr Asp Arg Thr Ile His Ser Ala Ala Gly Asp Pro Lys Ala Asn Gln Glu Ser Ile Glu Asn Asn Pro Phe Ala Pro Ser Gly Gln Gln Asp Ile 38.5 40 O Ser Ala Leu Tyr Thr Gly Leu Lys Val Thr Arg Gly Ile Trp Glu Ala Asp Phe Asn Leu Asn Tyr Thr Arg Asn Arg Ile Thr Gly Tyr Lys Pro Ala Cys Asp Ser Arg Val Ile Cys Val Pro Gln Gly Ser Tyr Asp Ile

, 5	Asp Gln	Asp 450	Lys	Glu	Gly	Gly	Phe 455	Asn	Pro	Ser	Val	Gln 460	Leu	Ser	Ala
10	Val Arg 465 480	Thr	Pro	Trp	Leu	Gln 470	Pro	Phe	Ile	Gly	Tyr 475	Ser	Lys	Ser	Met
15	Ala Ser	Pro	Asn	Ile	Gln 485	Glu	Met	Phe	Phe	Ser 490	Asn	Ser	Gly	Gly	Ala 495
20	Met Gly	Asn	Pro	Phe	Leu	Lys	Pro	Glu	Arg 505	Ala	Glu	Thr	Trp	Gln 510	Ala
. 25	Phe Arg	Asn	Ile 5 <u>1</u> 5	Asp	Thr	Arg	Asp _.	Leu 520	Leu	Val	Glu	Gln	Asp 525	Ala	Leu
,30	Phe Ser	Lys 530	Ala	Leu	Ala	Tyr	Arg 535	Ser	Arg	Ile	Gln	Asn 540	Tyr	Ile	Tyr .
35	Glu Glu 545 560	Ser 、	Tyr	Leu	Val [·]	Cys 550	Ser	Gly	Gly	Arg	Lys 555	Cys	Ser	Leu	·Pro
40	Val Asn	Ile	Gly	Asn	Gly 565	Trp	Glu	Gly	Ile	Ser 570	Asp	Glu	Tyr	Ser	Asp 575
45	Met Phe	Tyr	Ile	Tyr 580	Val	Asn	Ser	Ala	Ser 585	Asp	Val	Ile	Ala	Lys 590	Gly
50	Glu Ser	Leu	Glu	Met	Asp	Tyr.	Asp	Ala	Gly	Phe	Ala	Phe	Gly	Arg	Leu

			1/0/5/

595 600 605

Phe Ser Gln Gln Gln Thr Asp Gln Pro Thr Ser Ile Ala Ser Thr His 610 615 620

Phe Gly Ala Gly Asp Ile Thr Glu Leu Pro Arg Lys Tyr Met Thr
Leu
625
640
635

Asp Thr Gly Val Arg Phe Phe Asp Asn Ala Leu Thr Leu Gly Thr Ile 645 650

20 Ile Lys Tyr Thr Gly Lys Ala Arg Arg Leu Ser Pro Asp Phe Glu Gln 660 665 670

25 Asp Glu His Thr Gly Ala Ile Ile Lys Gln Asp Leu Pro Gln Ile Pro 675 680 685

30 Thr Ile Ile Asp Leu Tyr Gly Thr Tyr Glu Tyr Asn Arg Asn Leu Thr 690 695 700

35 Leu Lys Leu Ser Val Gln Asn Leu Met Asn Arg Asp Tyr Ser Glu
Ala
705 710 715

Leu Asn Lys Leu Asn Met Met Pro Gly Leu Gly Asp Glu Thr His Pro 725 730 735

Ala Asn Ser Ala Arg Gly Arg Thr Trp Ile Phe Gly Gly Asp Ile Arg
740 745 750

50 Phe

<210> 52 <211> 133 <212> PRT <213> Escherichia coli <400> Met Ser Ser Lys Thr Lys Cys Trp Leu Trp Met Leu Leu Val Ile Ser Glu Thr Ser Ala Thr Ser Thr Leu Lys Met Phe Asp Asn Ser Gly Met Thr Lys Thr Leu Leu Ala Leu Ile Val Val Leu Tyr Cys Ile Cys Tyr Tyr Ser Leu Ser Arg Ala Val Lys Asp Ile Pro Val Gly Leu Ala Tyr Ala Thr Trp Ser Gly Thr Gly Ile Leu Met Val Ser Thr Leu Gly Ile Leu Phe Tyr Gly Gln His Pro Asp Thr Ala Ala Ile Ile Gly Met Val Ile Ile Ala Ser Gly Ile Ile Ile Met Asn Leu Phe Ser Lys Met Gly Ser Glu Glu Ala Glu Glu Thr Pro Val Thr Asn Leu

Lys Lys Ile Ala Asn

<210>	53 <211>	286 <212>	PRT <213>	Escherichia coli	<400>
53			1		

Met Tyr Ile Lys Lys His Trp Ile Ala Leu Ser Ile Leu Leu Ile 5 Pro 1 5 10 15

Cys Ile Gly Asn Ala Gln Glu Ile Lys Ile Asp Glu Ser Trp Leu

10 His
20 25 30

Gln Ser Leu Asn Val Ile Gly Arg Thr Asp Ser Arg Phe Gly Pro
15 Arg
35 40 45

Leu Thr Asn Asp Leu Tyr Pro Glu Tyr Thr Val Ala Gly Arg Lys
20 Asp
50 55 60

Trp Phe Asp Phe Tyr Gly Tyr Val Asp Leu Pro Lys Phe Phe Gly
Val
65 70 75 80

Gly Ser His Tyr Asp Val Gly Ile Trp Asp Glu Gly Ser Pro Leu 30 Phe 85 90 95

Thr Glu Ile Glu Pro Arg Phe Ser Ile Asp Lys Leu Thr Gly Leu 35 Asn 100 105 110

Leu Ala Phe Gly Pro Phe Lys Glu Trp Phe Ile Ala Asn Asn Tyr 40 Val 115 120 125

Tyr Asp Met Gly Asp Asn Gln Ser Ser Arg Gln Ser Thr Trp Tyr 45 Met 130 · 135 140

Gly Leu Gly Thr Asp Ile Asp Thr Gly Leu Pro Ile Lys Leu Ser . $50\,$ Ala

145 150 155

160

5 Asn Ile Tyr Ala Lys Tyr Gln Trp Gln Asn Tyr Gly Ala Ala Asn Glu
165 170 175

10 Asn Glu Trp Asp Gly Tyr Arg Phe Lys Ile Lys Tyr Ser Ile Pro Leu 180 185 190

15 Thr Asn Leu Phe Gly Gly Arg Leu Val Tyr Asn Ser Phe Thr Asn Phe
195 200 205

20 Asp Phe Gly Ser Asp Leu Ala Asp Lys Ser His Asn Asn Lys Arg
Thr
210 215 220

25 Ser Asn Ala Ile Ala Ser Ser His Ile Leu Ser Leu Leu Tyr Glu His 225 230 235 240

30

Trp Lys Phe Ala Phe Thr Leu Arg Tyr Phe His Asn Gly Gly Gln
Trp

245

250

255

Asn Ala Gly Glu Lys Val Asn Phe Gly Asp Gly Pro Phe Glu Leu Lys

260 265 270

40
Asn Thr Gly Trp Gly Thr Tyr Thr Thr Ile Gly Tyr Gln Phe
275
280
285

50

45 <210> 54 <211> 172 <212> PRT <213> Escherichia coli <400> 54

Met Arg Ile Ala Pro Arg Thr Phe Phe Ala Ile Ser Ala Leu Ala Phe 1 5 10 15

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Ile Val Ala Ser Gly Phe Ser Phe Trp Arg Leu Ser Pro Ala Glu Asn Thr Gly Ile Met Ser Cys Ser Thr Lys Gly Ile Met Arg Phe Glu Asn Met Glu Lys Glu Asn Val Asn Gly Asn Ile His Phe Asn Phe Gly Ser Gln Gly Lys Gly Ser Met Val Leu Glu Gly Tyr Thr Asp Ser Ala Gly Trp Leu Tyr Leu Gln Arg Tyr Val Lys Phe Thr Tyr Thr Ser Lys Arg Val Ser Ala Thr Glu Arg His Tyr Arg Ile Ser Gln Trp Glu Ser Ser Ala Ser Ser Ile Asp Glu Ser Pro Asp Val Ile Phe Asp Tyr Met Arg Glu Met Ser Asp Ser His Asp Gly Leu Phe Leu Asn Ala Gln Lys Leu Asn Asp Lys Ala Ile Leu Leu Ser Ser Ile Asn Ser Pro Leu

Trp Ile Cys Thr Leu Lys Ser Gly Ser Lys Leu Asp

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<210> 55 <211> 182 <212> PRT <213> Escherichia coli <400> 55

Met Lys Ile Lys Val Ile Ala Leu Ala Thr Phe Val Ser Ala Val 5 Phe 1 5 10 15

Ala Gly Ser Ala Met Ala Tyr Asp Gly Thr Ile Thr Phe Thr Gly

10 Lys
20 25 30

Val Val Ala Gln Thr Cys Thr Val Asn Thr Ser Asp Lys Asp Leu

15 Ala

35 40 45

Val Thr Leu Pro Thr Val Ala Thr Ser Ser Leu Lys Asp Asn Ala
20 Ala
50 55 60

Thr Ser Gly Leu Thr Pro Phe Ala Ile Arg Leu Thr Gly Cys Ala
25 Thr
65 70 75 80

Gly Met Asn Ser Ala Gln Asn Val Lys Ala Tyr Phe Glu Pro Ser 30 Ser 85 90 95

Asn Ile Asp Leu Ala Thr His Asn Leu Lys Asn Thr Ala Thr Pro

Thr

100 105 110

Lys Ala Asp Asn Val Gln Ile Gln Leu Leu Asn Ser Asn Gly Thr 40 Ser 115 120 125

Thr Ile Leu Leu Gly Glu Ala Asp Asn Gly Gln Asp Val Gln Ser
45 Glu
130 135 140

Thr Ile Gly Ser Asp Gly Ser Ala Thr Leu Arg Tyr Met Ala Gln 50 Tyr

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145	150	155	•			

5 Tyr Ala Thr Gly Gln Ser Thr Ala Gly Asp Val Lys Ala Thr Val His 165 170 175

10 Tyr Thr Ile Ala Tyr Glu 180 \cdot

160

20

30

40

45

<210> 56 <211> 359 <212> PRT <213> Escherichia coli <400>
15 56

Met Lys Arg Ile Phe Phe Ile Pro Leu Phe Leu Ile Leu Leu Pro Lys

1 5 10 15

Leu Ala Val Ala Gly Pro Asp Asp Tyr Val Pro Ser Gln Ile Ala Val

20 25 30 25

Asn Thr Ser Thr Leu Pro Gly Val Val Ile Gly Pro Ala Asp Ala His

35
40
45

Thr Tyr Pro Arg Val Ile Gly Glu Leu Ala Gly Thr Ser Asn Gln Tyr

50 55 60 35

Val Phe Asn Gly Gly Ala Ile Ala Leu Met Arg Gly Lys Phe Thr Pro 65 70 75 80

Ala Leu Pro Lys Ile Gly Ser Ile Thr Val Tyr Phe Pro Ser Arg Lys 85 90 95

Gln Arg Asp Ser Ser Asp Phe Asp Ile Tyr Asp Ile Gly Val Ser Gly

100 105 110

50 .

	185/37
	185/3/

	Leu Leu	Gly	Ile		Ile	Gly	Met	Ala	Gly	Tyr	Trp	Pro	Ala 125	Thr	Pro
5	Val Asn	Pro	Ile	Asn	Ser	Ser		Ile	Tyr	Ile	Asp		Val	Gly	Ala
10	Thr Arg 145	Asn	Pro	Asn	Thr	Tyr 150	135 Asn	Gly	Ala	Thr	Ala 155			Gly	Ala
15	160												- (1)		
. 20	Leu Ile	Phe	Val	·Ala	Phe 165	Val	Ala	Thr	Gly	Arg 170	Lėu	Pro	Asn	Gly	Tyr 175
25	Thr Arg	Ile	Pro	Thr 180	Arg	Gln	Leu	Gly	Thr 185	Ile	Leu	Leu	Glu	Ala 190	Lys
30	Thr Gly	Ser	Leu 195	Asn.	Asn	Lys	Gly	Leu 200	Thr	Ala	Pro	Val	Met 205	Leu	Asn
35	Gly Asn	Arg 210	Ile	Gln	Val	Gln	Ser 215	Gln	Thr	Cys	Thr	Met 220	Gly	Gln	Lys
40	Tyr Leu 225 240	Val	Val	Pro	Leu	Asn 230	Thr	Val '	Tyr	Gln	Ser 235	Gln	Phe	Thr	Ser
45		Lys	Glu		Gln 245	Gly	Gly	Lys	Ile	Asp 250		His	Leu		Cys 255
50	Asp Val	Gly				Tyr	Ala		Leu 265			Ala			

186/3
100/3

Asn Arg Thr Asp Ile Leu Thr Leu Ser Ser Glu Ser Thr Ala Lys Gly

Phe Gly Ile Arg Leu Tyr Lys Asp Ser Asp Val Thr Ala Ile Ser

Gly Glu Asp Ser Pro Val Lys Gly Asn Gly Ser Gln Trp His Phe

Asp Tyr Arg Gly Glu Val Asn Pro His Ile Asn Leu Arg Ala Asn Tyr

Ile Lys Ile Ala Asp Ala Thr Thr Pro Gly Ser Val Lys Ala Ile

Ala

Thr Ile Thr Phe Ser Tyr Gln

<210> 57 <211> 844 <212> PRT <213> Escherichia coli <400>

Met Asn Ala Asn Asn Leu Ser Cys Leu Ile Tyr Cys Arg Cys Ser Leu

Leu Leu Phe Ala Ala Leu Gly Leu Thr Val Thr Asn His Ser Phe Ala

Ala Glu Glu Ala Glu Phe Asp Ser Glu Phe Leu His Leu Asp Lys Gly

Ile Asn Ala Ile Asp Ile Arg Arg Phe Ser His Gly Asn Pro Val Pro

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80

50 55 60

Glu Gly Arg Tyr Tyr Ser Asp Ile Tyr Val Asn Asn Val Trp Lys 5 Gly 65 70 75 Lys Ala Asp Leu Gln Tyr Leu Arg Thr Ala Asn Thr Gly Ala Pro 10 Thr 85 90 95 Leu Cys Leu Thr Pro Glu Leu Leu Ser Leu Ile Asp Leu Val Lys 15 Asp 100 105 110 Thr Met Ser Gly Asn Thr Ser Cys Phe Pro Ala Ser Thr Gly Leu 20 Ser 115 120 . 125 Ser Ala Arg Ile Asn Phe Asp Leu Ser Thr Leu Arg Leu Asn Ile 25 Glu 130 135 140 Ile Pro Gln Ala Leu Leu Asn Thr Arg Pro Arg Gly Tyr Ile Ser 30 Pro 145 150 155 160 Ala Gln Trp Gln Ser Gly Val Pro Ala Ala Phe Ile Asn Tyr Asp 35 Ala 165 170 175 Asn Tyr Tyr Gln Tyr Ser Ser Ser Gly Thr Ser Asn Glu Gln Thr 40 Tyr 180 185 190

45 Leu Gly Leu Lys Ala Gly Phe Asn Leu Trp Gly Trp Ala Leu Arg His 195 200 205

50 Arg Gly Ser Glu Ser Trp Asn Asn Ser Tyr Pro Ala Gly Tyr Gln Asn

210 215 220

Ile Glu Thr Ser Ile Met His Asp Leu Ala Pro Leu Arg Ala Gln . Phe Thr Leu Gly Asp Phe Tyr Thr Asn Gly Glu Leu Met Asp Ser Leu Leu Arg Gly Val Arg Leu Ala Ser Asp Glu Arg Met Leu Pro Gly Ser Leu Arg Gly Tyr Ala Pro Ala Val Arg Gly Ile Ala Asn Ser Asn Ala Lys Val Thr Ile Tyr Gln Asn Ala His Ile Leu Tyr Glu Thr Thr Val Pro Ala Gly Pro Phe Val Ile Asn Asp Leu Tyr Pro Ser Gly Tyr Ala Gly Asp Leu Leu Val Lys Ile Thr Glu Ser Asn Gly Gln Thr Arg Met Phe Thr Val Pro Phe Ala Ala Val Ala Gln Leu Ile Arg Pro Gly Phe Ser Arg Trp Gln Met Ser Val Gly Lys Tyr Arg Tyr Ala Asn Lys

355 360 365

	W	O 2 00	5/09782	3			189/370						PCT/EP2005/003972		
	Tyr Asn	Asn	Asp	Leu	Ile	Ala	Gln	Gly	Thr	Tyr	Gln	Tyr	Gly	Leu	Thr
		370					375					380			
5	_		1		_										•
	Asp Ala	TTE	Thr	Leu	Asn	Ser	GLy	Leu	Thr	Thr	Ala	Ser	Gly	Tyr	Thr
	385 400					390		,			395				
10				-											
	Gly Ser	Leu	Ala	Gly	Leu	Ala	Phe	Asn	Thr	Pro	Leu	Gly	Aļa	Ile	Ala

Asp Ile Thr Leu Ser Arg Thr Ala Phe Arg Tyr Ser Gly Val Thr . Arg

410

415

420 425 430

405

15

20

25

30

35

40

Lys Gly Tyr Ser Leu His Ser Ser Tyr Ser Ile Asn Ile Pro Ala Ser 435 440 445

Asn Thr Asn Ile Thr Leu Ala Ala Tyr Arg Tyr Ser Ser Lys Asp Phe

450 455 460

Tyr His Leu Lys Asp Ala Leu Ser Ala Asn His Asn Ala Phe Ile Asp 465 470 475 480

Asp Val Ser Val Lys Ser Thr Ala Phe Tyr Arg Pro Arg Asn Gln Phe
485 490 495

Gln Ile Ser Ile Asn Gln Glu Leu Gly Glu Lys Trp Gly Gly Met Tyr
45 500 505 510

Leu Thr Gly Thr Thr Tyr Asn Tyr Trp Gly His Lys Gly Ser Arg
Asn
50 515 520 525

Glu Gln	Tyr	Gln	Ile	Gly	Tyr	Şer	Asn	Phe	Trp	Lys	Gln	Leu	Gly	Tyr
	530					535					540			

Ile Gly Leu Ser Gln Ser Arg Asp Asn Glu Gln Gln Arg Arg Asp Asp

Arg Phe Tyr Ile Asn Phe Thr Leu Pro Leu Gly Gly Ser Val Gln

Pro Val Phe Ser Thr Val Leu Asn Tyr Ser Lys Glu Glu Lys Asn

Ile Gln Thr Ser Ile Ser Gly Thr Gly Glu Asp Asn Gln Phe Ser

Tyr Gly Ile Ser Gly Asn Ser Gln Glu Asn Gly Pro Ser Gly Tyr

Met Asn Gly Gly Tyr Arg Ser Pro Tyr Val Asn Ile Thr Thr Val

Gly His Asp Thr Gln Asn Asn Gln Arg Ser Phe Gly Ala Ser Gly 655 .

Ala Val Val Ala His Pro Tyr Gly Val Thr Leu Ser Asn Asp Leu Ser

Asp Thr Phe Ala Ile Ile His Ala Glu Gly Ala Gln Gly Ala Val Ile

	Asn Val	Asn	Ala	Ser	Gly	Ser	Arg	Leu	Asp	Phe	Trp	Gly	Asn	Ģly	Val
5		690					695					700			
	Pro Ser	Tyr	Val	Thr	Pro	Tyr	Glu	Lys	Asn	Gln	Ile	Ser	Ile	Asp	Pro
10	705 720					710					715				
1.7		Leu	Asp	Leu	Asn	Val	Glu	Leu	Ser	Ala	Thr	Glu	Gln	Glu	Ile
15 .	Ile				725			٠		730					735
20		Arg	Ala	Asn	Ser	Ala	Thr	Leu	Val	Lys	Phe	Asp	Thr	Lys	Thr
20	Gly			740					745					750	
25		Ser	Leu	Leu	Phe	Asp	Ile	Arg	Met	Ser	Thr	Gly	Asn	Pro	Pro
25	Pro		755					760				•	765		,
30		Ala	Ser	Glu	Val	Leu	Asp	Glu	His	Gly	Gln	Leu	Ala	Gly	Tyr
30	Val	770				٠	775					780			
35	Ala His	Gln	Ala	Gly	Lys	Val	Phe	Thr	Arg	Gly	Leu	Pro	Glu	Lys	Gly
<i>33</i>	785 800					790					795				
40		Ser	Val	Val	Trp	Gly	Pro	Asp	Asn	Lys.	Asp	Arg	Cys	Ser	Phe
	Val				805					810					815
45		His	Val	Ala	His	Asn	Lys	Asp	Asp	Met	Gln	Ser	Gln	Leu	Val
	Pro		-	820			٠		825					830	

50 . Val Leu Cys Ile Gln His Pro Asn Gln Glu Lys Thr 835

192/370

<210> 58 <211> 277 <212> PRT <213> Escherichia coli <400> Met Val Lys Cys His Thr Leu Ile Asn Arg Arg Asn Lys Cys Leu Leu 1 . Ile Val Phe Ile Val Leu Ile Gly Trp Ile Ile Phe Arg Pro Lys Ala Tyr Thr Tyr Ser Leu Asn Asp Lys Glu Lys Glu Met Leu Ile Met Leu Ser Gln His Pro Glu Thr Arg Tyr Phe Gly Phe Tyr Ser Ile Glu Leu Pro Ala Asp Tyr Lys Pro Thr Gly Met Val Met Phe Ile Gln Gly Ser Ala Met Ile Pro Val Glu Thr Lys Leu Gln Tyr Tyr Pro Phe Leu Gln Tyr Met Thr Arg Tyr Glu Ala Glu Leu Lys Asn Thr Ser Ala Leu Asp Pro Leu Asp Thr Pro Tyr Leu Lys Gln Val His Pro Leu Ser Pro Met Asn Gly Val Ile Phe Glu Arg Met Lys Ala Lys Tyr Thr

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	402/250	

Asp Phe Ala Arg Val Leu Asp Ala Trp Lys Trp Glu Asn Gly Val Thr
145 150 155

5

Phe Ser Val Lys Ile Glu Ala Lys Asp Gly Arg Ala Thr Arg Tyr Asp

165 170 175

10

Gly Ile Ser Lys Ile Ala Glu Tyr Ser Tyr Gly Tyr Asn Ile Pro Glu
180 185 190

15

Lys Lys Val Gln Leu Leu Thr Ile Leu Ser Gly Leu Gln Pxo Arg Ala 195 200 205

20

Asp Asn Gln Pro Pro Ser Glu Asn Lys Leu Ala Ile Gln Tyr Ala Gln
210 215 220

25

Val Asp Ala Ser Leu Leu Gly Glu Tyr Glu Leu Ser Val Asp Tyr Lys
230 235

30 240

Asn Ser Asn Asn Ile Lys Ile Ser Leu Gln Thr Asp Asn Asn Ser Tyr

35 245 250 255

Ile Asp Ser Leu Leu Asp Ile Arg Tyr Pro Ser Asn Gly Asn Arg Ala 40 260 265 270

Trp Tyr Asn Ser Ile 275

45

<210> 59 <211> 366 <212> PRT <213> Escherichia coli <400> 59

50 Met Leu Pro Glu Pro Val Tyr Arg Arg Trp Ile Ile Leu Leu Ile Ser

	V	VO 200	5/0978	23			194/370					PCT	PCT/EP2005/003972			
	1				5					10	•				15	
5	Met Ala	Leu	Thr	Val	Gly	Thr	Leu	Phe	Ile	Leu	Ser	Val	Trp	Asn	Ser	
	711.0			20					25	•				30		
10	Thr Leu	Tyr	Trp	Asp	Ile	Phe	Ile	Tyr	Gly	Val	Leu	Pro	Met	Leu	Phe	-
			35					40					4 5			
15	Trp Val	Leu	Cys	Leu	Phe	Gly	Ile	Alą	Leu	Asn	Lys	Tyr	Glu	Gln	Ser	
,	-	50					55					60		et i		
20	Ala Leu	Ala	Cys	Ile	Ser	Trp	Glu	Ser	Glu	Arg	Gln	Gln	Val	Lys	Gln	
20	65					70					75					80
25	Trp Leu	Gln	His	Trp	Ser	Gln	.Lys	Gln	Leu	Ala	Ile	Val	Gly	Asn	Val	
23	шец				85					90					95	
	Phe	Thr	Pro	Glu	Glu	Lys	Gly	Met	Ser	Val	Leu	Leu	Gly	Pro	Gln	,

Phe Thr Pro Glu Glu Lys Gly Met Ser Val Leu Leu Gly Pro Gln
Glu
100 105 110

Arg Tyr Ser Leu Ser Ser Ile Phe His Asp Ile His Gln Gln Leu 40 Thr 130 135 140

Gln Gln Phe Pro Asp Tyr Arg His Tyr Leu His Thr Ile Tyr Val 45 Leu 145 150 155

50 Gln Pro Glu Lys Trp Arg Gly Glu Thr Val Arg Gln Ala Ile Phe His

165 170 175

- Gln Trp Asp Leu Val Pro Glu Arg Thr Asn Thr Leu Asn Gln Ile 5 Gln 180 185 190
- Ser Leu Tyr Asp Glu Arg Phe Asp Gly Leu Ile Leu Val Val Cys 10 Leu 195 200 205
- Gln Asn Trp Pro Glu Asn Lys Pro Glu Asp Thr Ser Glu Leu Val 15 Ser 210 215 220
- Ala Gln Leu Ile Ser Ser Ser Phe Val Arg Gln His Gln Ile
 Pro
 225 230 235
 240
- Val Ile Ala Gly Leu Gly Arg Val Met Pro Leu Glu Pro Glu Glu Leu
 245
 250
 255
- 30 Glu His Asn Leu Asp Val Leu Phe Glu Tyr Asn Gln Leu Asp Asn Lys 260 265 270
- 35 Gln Leu Gln His Val Trp Val Ser Gly Leu Asp Glu Gly Thr Ile Glu 275 280 285
- 40 Asn Leu Met Gln Tyr Ala Glu Gln His Gln Trp Ser Leu Pro Lys Lys 290 295 300
- 45 Arg Pro Leu His Met Ile Asp His Ser Phe Gly Pro Thr Gly Glu Phe 305 310 315

W	O 2005	5/09782	23		196/370							PCT/EP2005/003972				
Ile Thr	Phe	Pro	Val	Ser	Leu	Ala	Met	Leu	Ser	Glu	Ala	Ala	Lys	Glu		

330

335

Glu Gln Asn His Leu Ile Ile Tyr Gln Ser Ala Gln Tyr Ala Gln Lys

340
345
350

Lys Ser Leu Cys Leu Ile Thr Arg Lys Leu Tyr Leu Arg Thr 355 360 365

325

15 <210> 60 <211> 260 <212> PRT <213> Escherichia coli <400>

Met Leu Asn Arg Lys Leu Asn Ile Arg Leu Arg His Ser Leu Asn Ser 20 1 5 10 15

His Cys Ile Pro Ser Ile Ile Ile Asn Asn Thr Val Arg Ser Phe Gln 25 20 25 30

Arg Ser Val Met Asn Thr Arg Ala Leu Phe Pro Leu Leu Phe Thr Val

30 35 40 45

Ala Ser Phe Ser Ala Ser Ala Gly Asn Trp Ala Val Lys Asn Gly Trp 35 55 60

Cys Gln Thr Met Thr Glu Asp Gly Gln Ala Leu Val Met Leu Lys Asn 40 65 70 75 80

Gly Thr Ile Gly Ile Thr Gly Leu Met Gln Gly Cys Pro Asn Gly Val
45 85 90 95

Gln Thr Leu Leu Gly Ser Arg Ile Ser Ile Asn Gly Asn Leu Ile Pro 50 100 105 110

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	197/370	

Thr Ser Gln Met Cys Asn Gln Gln Thr Gly Phe Arg Ala Val Glu Val Glu Ile Gly Gln Ala Pro Glu Met Val Lys Lys Ala Val His Ser . 10 Ala Glu Arg Asp Val Ser Val Leu Gln Ala Phe Gly Val Arg Met Glu Phe Thr Arg Gly Asp Met Leu Lys Val Cys Pro Lys Phe Val Thr Ser Leu Ala Gly Phe Ser Pro Lys Gln Thr Thr Thr Ile Asn Lys Asp Val Leu Gln Ala Ala Arg Gln Ala Tyr Ala Arg Glu Tyr Asp Glu Thr Thr Glu Thr Ala Asp Phe Gly Ser Tyr Glu Val Lys Gly Asn Lys Val Glu Phe Glu Val Phe Asn Pro Glu Asp Arg Ala Tyr Asp Lys Val Thr Val Thr Val Gly Ala Asp Gly Asn Ala Thr Gly Ala Ser Val Glu Phe Ile Gly Lys

198/37/0

<210> 61 <211> 385 <212> PRT <213> Escherichia coli <400> 61

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- 10 Ser Leu Thr Ser Leu Leu Pro Asp Ile Arg Lys Met Leu Leu Val Thr 20 25 30
- 15 Asp Arg Asn Ile Ala Gln Leu Asp Gly Val Gln Gln Ile Arg Ala Leu 35 40 45
- 20 Leu Glu Lys His Cys Pro Gln Val Asn Val Ile Asp Asn Val Pro Ala 50 55 60
- 25 Glu Pro Thr His His Asp Val Arg Gln Leu Met Asp Ala Pro Gly Asp 65 70 75 80
- 30 Ala Ser Phe Asp Val Val Val Gly Ile Gly Gly Gly Ser Val Leu Asp 85 90 95
- Val Ala Lys Leu Leu Ser Val Leu Cys His Pro Gln Ser Pro Gly Leu
 100
 105
 110
- 40 Asp Ala Leu Leu Ala Gly Glu Lys Pro Thr Gln Arg Val Gln Ser Trp
 115 120 125
- 45 Leu Ile Pro Thr Thr Ala Gly Thr Gly Ser Glu Ala Thr Pro Asn Ala
 130 135 140
- 50 Ile Leu Ala Ile Pro Glu Gln Ser Thr Lys Val Gly Ile Ile Ser Gln

Ĭ

	V	VO 200	J3/UY /8	523				199/3	3 70				PCI	EP200	5/0039
	145 160					150					155				
5	Val Ser	Leu	Leu	Pro	Asp	Tyr	Val	Ala	Leu	Phe	Pro	Glu	Leu	Thr	Thr
					165					170					175
10		Pro	Ala	His	Ile	Ala	Ala	Ser	Thr	Gly	Ile	Asp	Ala	Leu	Cys
	His			180					185					190	
15		Leu	Glu	Cys	Phe	Thr	Ala	Thr	Val	Ala	Asn	Pro	Val	Ser	Asp
	Asn		195	*				200					205		
20	Ala Ala	Ala	Leu	Thr	Gly	Leu	Ser	Ļys	Leu	Phe	Arg	His	Ile	Gln	Pro
	AIG	210	3	*			215					220			
25	Val Ala	Asn	Asp	Pro	Gln	Asp	Leu	Arg	Ala	Lys	Leu	Glu	Met	Leu	Trp
	225 240					230					235				
30	Ser	Tur	Tur	Gly	Glv	Val	Ala	Tle	Thr	His	Ala	Glv	Thr	His	T. 2 11
	Val	-1-		J—1	245					250		1		1120	255
35															200
	His Gly	Ala	Leu	Ser	Tyr	Pro	Leu	Gly	Gly	Lys	Tyr	His	Leu	Pro	His
	4			260					265					270	
40		Ala	Asn	Ala	Ile.	Lèu	·Leu	Ala	Pro	Cys	Met	Ala	Phe	Val	Arg
Pro		275					280					285			

45 Trp Ala Val Glu Lys Phe Ala Arg Val Trp Asp Cys Ile Pro Asp Ala 290 295 300

	V	VO 200	5/0978	23				200/3	370				PCT/	EP200	5/003972
5	Glu Trp 305 320	Thr	Ala	Leu	Ser	Ala 310	Glu			Ser	His		Leu	Val	Thr
	Leu Ala	Gln	Ala	Leu	Val 325	Asn	Gln	Leu	Lys	Leu 330	Pro	Asn	Asn	Leu	
10					323					330					335
	Leu Leu	Gly	Val	Pro	Pro	Glu	Asp	Ile	Ala	Ser	Leu	Ser	Glu	Ala	Ala
15	•			340					345					350	
	Asn Gln	Val	Lys	Arg	Leu	Met	Asn	Asn	Val	Pro	Cys	Gln	Ile	Asp	Leu
20			355					360	*				365		
	Asp Lys	Val	Gln	Ala	Ile	Tyr	Gln	Thr	Leu	Phe	Pro	Gln	His	Pro	Phe
25	-	370	•				375					380			
	Glu 385	•													,
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	1		<i>11</i> *		5					10 .					15
40	Leu Cys	Phe	Leu	Phe	Phe	Phe	Val	Val	Ser	Ala	Ile	Thr	Thr	Ile	Ala
	~1°			20					25					30	

Gly Tyr Thr Glu Lys Asn Ala Thr Gly Asn Val Leu Leu Phe
45 Leu
35 40 45

Leu Leu Leu Ala His Arg Asn Thr Leu Thr Ser Ile Thr Ala 50 Leu 50 55 60

														,		
5	Leu Tyr 65	Phe	Leu	Phe	Cys	Cys 7.0	Ala	Leu	Tyr	Ala	.Pro 75	Ala	Gly	Met	Thr	80
10	Gly Thr	Lys	Ile	Asn	Asn 85	Ser	Phe	Ile	Val	Ala 90	Leu	Leu	Gln	Thr	Thr 95	
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25	Leu Tyr	Phe	Ile	Phe 20	Phe	Leu	Ser	Ser	Ala 25	Phe	Val	Ser	Phe	Gly 30	Cys	
30	Ala Trp	Ile	Tyr 35	Glu	Leu	Phe	Leu	Trp	Asn	Asp	Ile	Ile	Val	Tyr	Ser	
35	Gly Ser	Tyr 50	Ile		Ile	Val	Phe 55	Leu	Pro	Phe	Thr	Leu 60	Tyr	Val	Met	
40	Phe Val 65	Glu	Ile	Leu	Phe	Phe	Ala	Ile	Ser	Gly	Arg 75	Arg	Leu	Ser	Lys	80
45	Thr Ser	Met	Val	Arg	Leu 85	Trp	Leu	Ile	Ile	Lys 90	Ile	Ile	Ile	Ala	Phe 95	
50	Ile	Cys	Ala	Val	Leu	Ile	Phe	Ser	Ser	Ile	Tyr	Lys	Lys	Glu	Leu	

Leu

100 105 110

Ser Arg Asn Tyr Ile Ala Cys Ser Gly Ile Pro Ser Gly Trp Met 5 Pro 115 120 125

Gly Leu Ala Thr Lys Tyr Val Lys Glu Lys Ser Leu Cys Glu Lys

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130 135 140

Gly Asn Asn . 15 145

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Met Phe Pro Ile Arg Phe Lys Arg Pro Ala Leu Leu Cys Met Ala Met
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Leu Thr Val Val Leu Ser Gly Cys Gly Leu Ile Gln Lys Val Val Asp
20 25 30

30
Glu Ser Lys Ser Val Ala Ser Ala Val Phe Tyr Lys Gln Ile Lys Ile
35
40
45

Leu His Leu Asp Phe Phe Ser Arg Ser Ala Leu Asn Thr Asp Ala Glu
50 55 60

Asp Thr Pro Leu Ser Thr Met Val His Val Trp Gln Leu Lys Thr Arg 65 70 75 80

Glu Asp Phe Asp Lys Ala Asp Tyr Asp Thr Leu Phe Met Gln Glu Glu

85 90 95

	V	VO 200	05/0978	323				203/3	370				PCT/	EP2005	5/003972
	Lys Lys	Thr	Leu	Glu	Lys	Asp	Val	Leu	Ala	Lys	His	Thr	Val	Trp	Val
	-1-			100					105					110	
5	Pro Gln	Glu	Gly	Thr	Ala	Ser	Leu	Asn	Val	Pro	Leu	Asp	Lys	Glu	Thr
			115					120				n	125		. ,
10	Phe Asp	Val	Ala	Ile	Ile	Gly	Gln	Phe	Tyr	His	Pro	Asp	Glu	Lys	Ser
	1150	130					135					140			
15	Ser Pro	Trp	Arg	Leu	Val	Ile	Lys	Arg	Asp	Glu	Leu	Glu	Ala	Asp	Lys
20	145 160		ø			150		*			155				•
20	Arg	Ser	Ile	Glu	Leu	Met	Arg	Ser	Asp	Leu	Arg	Leu	Leu	Pro	Leu
25	Lys				165					170					175
	Asp	Lys					•							,	
30												•			•
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40	Ile Gln	Thr	Leu	Cys	Gly	Gly	Gly	Tyr	Tyr	Met	Tyr	Arg	Gln	Glu	Tyr
				20					25					30	
45	Met Pro	Val	Val	Thr	Val	Pro	Thr	Ala	Asp	Ala	Asn	Asp	Pro	Asn	Trp
			35					40					45		
50	Asn	Lys	Arg	Ile	Gln	Phe	Asp	Thr	Ser	Glu	Trp	Leu	Gln	Gln	Leu

Gln

5	Tyr Pro 65	Ile	Lys	Ile	Asp	Asp	His	Tyr	Ile	Leu	Asn 75	Thr	Gln	Tyr	Thr	80
10	Ile Ala	Ala	Asn	Leu	Asp 85	Asp	Phe	Gly	Ile	Thr	Leu	Lys	Leu	Gln	Asn 95	
15	Leu Glu	Asn	Gly	Ser	Asp	Lys	Arg	Leu	Pro 105	Ala	Leu	Tyr	Gly	Leu 110	Ala	
20	Met Cys	Asp	Ala 115	Gln	Lys	Phe	Lys	Asp 120		Met	Arg	Gly	Lys 125	Ile	Lys	
25	Glu Asn	Tyr 130	Leu	Arg	Thr	Thr	Phe	Asp	Ala	Glu	Thr	Leu 140	Lys	Pro	Val	
30.	Asp Phe 145 160	Tyr	Phe	Leu	Ile	Ser	Phe	Thr	Tyr	Lys	Asp	Lys	Trp	Tyr	Glu	
35	Glu Leu	Thr	Glu	Arg	Lys 165	Ile	Ser	Lys	Thr	Ser 170	Asp	Asp	Gly	Tyr	Phe	,
40	Trp Thr	Ala	Phe	Asp 180	Asn	Thr	Val	His	Glu 185	Ala	Gly	Tyr	Trp	His 190	Asn	
45	Asp Val	Pro	Ala 195	Ala	Tyr	Ser	Tyr	Arg 200	Asp	Tyr	Gln	Asn	Gly 205	Lys	Ala	
50	Lys					·										

	205/370															
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•	ALG			20		•			25					30		
15		Met	Asn	Ile	Phe	Thr	Leu	Ser	Lys	Ala	Pro	Leu	Tyr	Leu	Leu	
	Ile		35					40					45			
20	Ser Arg	Leu	Phe	Leu	Pro	Thr	Met	Ala	Met	Ala	Ile	Asp	Pro	Pro	Gĺu	
	ALG	50					55					60				
25	Glu Asp	Leu	Ser	Arg	Phe	Ala	Leu	Lys	Thr	Asn	Tyr	Leu	Gln	Ser	Pro	
	65					7.0					75 ·	,				80
30	Glu Phe	Gly	Val	Tyr	Glu	Leu	Ala	Phe	Asp	Asn	Ala	Ser	Lys	Lys	Val	
		•			85					90					95	
35	Ala Leu	Ala	Val	Thr	Asp	Arg	Val	Asn	Arg	Glu	Ala	Asn	Lys	Gly	Tyr	
	пси			100					105					110		
40	Tyr Met	Ser	Phe	Asn	Ser	Asp	Ser	Leu	Lys	Va l	Glu	Asn	Lys	Tyr	Thr	
	,		115					120					125	٠		
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	มอน	130					135					140	-			

Tyr Ile Gly His Thr Gln Ser Ala Ser Leu Arg Ile Ser Met Phe Asp

145 150 155 160

10 Ala Ala Asn Ala Ala Asp Ser Arg Phe Glu His Phe Arg His Met Val 180 185 · 190

15 Tyr Ser Gln Asp Ser Asp Thr Leu Phe Val Ser Tyr Ser Asn Met Leu
195 200 205

20 Lys Thr Ala Glu Gly Met Lys Pro Leu His Lys Leu Leu Met Leu Asp
210 215 220

25 Gly Thr Thr Leu Ala Leu Lys Gly Glu Val Lys Asp Ala Tyr Lys Gly 225 230 235

Thr Ala Tyr Gly Leu Thr Met Asp Glu Lys Thr Gln Lys Ile Tyr Val

245
250
255

35
Gly Gly Arg Asp Tyr Ile Asn Glu Ile Asp Ala Lys Asn Gln Thr
Leu
260
265
270

Leu Arg Thr Ile Pro Leu Lys Asp Pro Arg Pro Gln Ile Thr Ser Val . 275 280 285

45
Gln Asn Leu Ala Val Asp Ser Ala Ser Asp Arg Ala Phe Val Val
Val
290
295
300

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Phe Asp His Asp Asp Arg Ser Gly Thr Lys Asp Gly Leu Tyr Ile Phe 305 310 315 320

5

Asp Leu Arg Asp Gly Lys Gln Leu Gly Tyr Val His Thr Gly Ala 325 330 335

10

Ala Asn Ala Val Lys Tyr Asn Pro Lys Tyr Asn Glu Leu Tyr Val 340 345 350

15

Asn Phe Thr Ser Gly Thr Ile Ser Val Val Asp Ala Thr Lys Tyr Ser 355 360 . 365

20

Ile Thr Arg Glu Phe Asn Met Pro Val Tyr Pro Asn Gln Met Val Leu 370 375 380

25 .

Ser Asp Asp Met Asp Thr Leu Tyr Ile Gly Ile Lys Glu Gly Phe Asn 385 390 395 400

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Arg Asp Trp Asp Pro Asp Val Phe Val Glu Gly Ala Lys Glu Arg Ile 405 410 415

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ccagggaatg tggatgtttc tttgggtaat ctgtatgtat cagactttcc 50 caatgcagga 180

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10	gctggcggta atatcacaat	tcaagattga 360	aattcaggac	agggatggaa	gtaatgcatc
	ggtatgttca taaagcccgt	agacgcttaa 420	tgtacaaaat	aataatgcaa	cctttaatct
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	ccgcgcggct taagttcaac	atattgaagg 540	cggcgtcagt	agccgcgaca	gttatcgaag

ctgageggee ceatteagga tggeetgetg taeggeageg teaceetgtt aegeeaggtt 600

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	aagggccgta gcgctgcact	agctgtcgat 840 ·	cagcgatggt	tcaccagacc	cgtacatgcg
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	65					70					75					80
25	Gly Val	Gly	Leu	Ser	Ala	His	Leu	Ser	Thr	Asp	Ser	Pro	Val	Arg	Gln	
					85					90					95	
30	Ser Ala	Glu	Gly	Leu	Tyr	Leu	His	Val	Pro	Asp	Glu	Glu	Gln	Leu	Val	
	1114			100		•			105					110		
35		Val	Glu	Arg	Leu	Leu	Thr	Leu	Arg	Pro	Ala	Trp	Ala	Ser	Gln	
	Leu		115					120					125			
40																
	Ala Val	Val	Ala	Tyr	Thr	Ile	Met	Pro	Gly	Ile	His	Arg	Asp	Val	Ala	
		130					135					140			•	
45	Leu	Ala	Gly	Arg	Leu	Arg	Arg	Phe	Ala	His	Ser	Met	Ala	Thr	Val	•
50	Arg 145 160					150		ı			155					

WO 2005/097823	PCT/EP2005/003972

	V	VO 200	05/0978	23		240/250								PCT/EP2005/0039			
								310/3	370								
	Arg Ser	Arg	Ala	Gly	Val	Asn	Val	Pro	Trp	Leu	Leu	Trp	Ser	Gly	Leu		
					165					170			-		175		
5	Glv	Ser	Pro	Leu	Pro	Glu	Ara	Ala	Ser	Ser	Pro	Tro	Phe	Tla	Cve		
	Thr			180		O O	9	1110	185		110	- L	1110		Cyb		
10	,			100	÷				100					190	•		
10		Gly	Glu	Val	Gln	Val	Ala	Thr	Ser	Thr	Glu	Thr	Thr	Met	Pro		
	Ala		195					200				-	205				
15																	
	Gln Cys	Trp	Ile	Ala	Gln	Ser	Gly	Val	Gln	Glu	Arg	Ser	Gln	Arg	Leu		
		210					215					220					
20	Tvr	Leu	Leu	Lvs	Ala	Glu	Ser	Leu	Met	Gln	Trp	Leu	Asn	Leu	Asn		
	Val 225					230					235						
25	240					200					200						
40	T.011	Thr.	Ala	T.All	Δen	Glv	Pro	Clu	7\] ¬	T 17.0	Cvc.	Dro	Dro	Tou	7.1.		
	Met	1111	7114	шси	245	Ory	110	Grá	AIA		Суз	FIO	,F±O	пеп			
30					245					250					255		
		Vаl	Gly	Leu	Val	Pro	Ser	Leu	Pro	Ala	Val	Asp	Asn	Asn	Leu		
2.5	Trp			260					265		•	•		270			
35																	
	Gln Asp	Leu	Trp	Ile	Thr	Ala	Arg	Thr	Gly	Leu	Thr	Pro	Asp	Ile	Ala		
40			275 _.					280					285				
	Thr Gln	Gly	Thr	Asp	Asp	Ala	Leu	Pro	Phe	Pro	Asp	Ala	Ľeu	Leu	Arg		
45	GTII	290			·		295			:		300					
4		_															

Leu Pro Arg Gln Ser Gly Phe Thr Pro Leu Arg Arg Ala Cys Val Thr

															-
	Met Ser	Leu	Gly	Val	Thr	Thr	Val	Ala	Gly	Ile	Ala	Ala	Leu	Cys	Leu
5	per				325					330					335
	Ala His	Thr	Ala	Asn	Arg	Gln	Leu	Leu	Arg	Gln	Val	Gly	Asp	Asp	Leu
10				340					345					350	
	Arg His	Phe	Tyr	Ala	Val	Pro	Val	Glu	Glu	Phe	Ile	Thr	Lys	Ala	Arg
15			355					360					365		
٠	Leu Arg	Ser	Val	Leu	Lys	Asp	Asp	Ala	Thr	Met	Leu	Asp	Gly	Tyr	Tyr
20		370					375					380			
	Glu Arg	Gly	Glu	Pro	Leu	Arg	Leu	Gly	Leu	Gly	Leu	Tyr	Pro	Gly	Glu
25	385 400					390					395				
	Tle	Ara	Gln	Pro	Val	Len	Ara	Ala	Tle	Ara	Asp	Tro	Ara	Pro	Pro
30	Glu	9			405	,	,			410	F		9		415
	Gln	T.VS	Met	Glu	Val	Thr	Δla	Ser	T.e.ii	Gln	U = 1	Gln	Thr	Val	7\ 200
35	Leu	<i></i> 40	1100	420	var	****	TIEG	SCI	425	OTII	Val	0111	1111	430	Arg
	7\	Sor	Mot	502	Leu	Pho	7\ 5.70	W-1	Clv	Cln	7\ 1 ->	71 20 00	T 012	T	7. ~ ~
40	Gly.	Ser	435	per	пец	rne	ASP	440	GīĀ	GTII	ALA	ALG	445	тЛ2	ASP
	G .	ma 1	-	T 7	-	** 7	71		-	** 7	7 0	1	-		_
45	Pro	450	ГЛЗ	Val	Leu	Val	455	Ala	Leu	Val	Asn	11e 460	Arg	Ala	Lys
50	Gly Glu 465	Trp	Leu	Ile	Leu	Val 470	Ala	Gly	Tyr	Thr	Asp 475	Ala	Thr	Gly	Asp

5	Lys Asn		Asn	Gln	Gln 485	Leu	Ser	Leu	Arg	Arg 490	Ala	Glu	Ala	Val	Arg 495
10	Trp Gln	Met	Leu	Gln 500	Thr	Ser	Asp	Ile	Pro 505	Ala	Thr	Cys	Phe	Ala 510	Val
15	Gly Gly	Leu	Gly 515	Glu	Ser	Gln	Pro	Ala 520	Ala	Thr	Asn	Asp	Thr 525	Pro	Gln
20	Arg Asp	Ala 530	Val	Asn	Arg	Arg	Val 535	Glu	Ile	Ser	Leu	Val 540	Pro	Arg	Ser
25	Ala 545	Cys	Gln	Asp	Val	Lys 550									
	<210)> 3	L35 <	(211)	> 10	94 <2	212>	PRT	7 <21	3>	Esch	nerio	rhia	coli	
	<400		L35					2.202	-	.57	1001	10110)11±U	COTI	-
30)> 1	L35				Trp								
30	Met Leu 1)> 1	Lys	Ser	Thr 5	·Phe		Arg	Ala	Leu 10	Ala	Leu	Thr	Ala	Thr 15
	Met Leu 1 Ile Pro	Ile Leu	Lys Thr	Ser Gly 20	Thr 5 Cys	Phe Ser	Trp	Arg Ser	Ala Gln 25	Leu 10 Pro	Ala Glu	Leu Gln	Thr Glu	Ala Gly 30	Thr 15 Arg
35	Met Leu 1 Ile Pro Gln Gly	Ile Leu Ala	Lys Thr Trp 35	Ser Gly 20 Leu	Thr 5 Cys Gln	Phe Ser	Trp	Arg Ser Thr 40	Ala Gln 25 Leu	Leu 10 Pro	Ala Glu Thr	Leu Gln Leu	Thr . Glu Pro 45	Ala Gly 30 Ala	Thr 15 Arg

313/37/0

Ile Thr Leu Ala Gly Leu Ser Ser Val Gly Ile Arg Leu Phe Leu Val Thr Tyr Asp Ala Lys Gly Leu Arg Ala Glu Gln Ser Ile Val Val Pro Gln Leu Pro Pro Ala Ser Gln Val Leu Ala Asp Val Met Leu Ser His Trp Pro Ile Ser Ala Trp Gln Pro Gln Leu Pro Thr Gly Trp Thr Arg Asp Asn Gly Asp Lys Arg Glu Leu Arg Asn Ala Ser Gly Lys Leu Val Thr Glu Ile Thr Tyr Leu Asn Arg Gln Gly Lys Arg Val Pro Ile Ser Ile Glu Gln His Val Phe Lys Tyr His Ile Thr Ile Gln Tyr Leu Gly Asp

<210> 136 <211> 129 <212> PRT <213> Escherichia coli
45 <400> 136

Met Lys Arg Tyr Ile Lys Trp Phe Ala Ile Thr Ile Phe Ile Ser Met 5 10 15

	W	O 2005	/09782	3				314/3′	70				PCT/E	P2005	003972	2
	Leu Thr	Ser	Ala	Cys	Val	Arg	Thr	Ala	Pro	Val	Gln	Gln	Ile	Ser	Thr	
	4 4 4 4 4			20					25					30		
5	Val	Ser	Val	Gly	His	Thr	Gln	Glu	Gln	Val	Lys	Asn	Ala	Ile	T.e.i	
	Lys		35					40			-		45		200	
10																
	Ala Val		Ala	Gln	Arg	Lys		Ile	Met	Thr	Gln	Val	Ser	Pro ·	Gly	
1.5		50					55					60				
15	Ile	Lys	Ala	Arg	Tyr	Gln	Thr	Arg	Asn	His	Val	Ala	Glu	Val	Arg	
	Ile 65					70					75					80.
20	Thr	ጥላፖ	Ψhr	7.15	Thr	ጥ፣፣ጽ	Ф	7.00	тіс	T	П	7	G	G -	_	
	Asn	т у т.	T111	лта	85	туг	тут	ASII	тте	луs 90	Tyr	Asp	ser	ser	٠	•
25					00					90		•			95	
	Leu Val	Gln	Ala	Ser	Asp	Gly	Lys	Ile	His	Lys	Asn	Tyr	Asn	Arg	Trp	
	•			100					105					110		
30	Arg	Asn	Leu	Asp	Lys	Asp	Ile	Gln	Val	Asn	Leu	Ser	Thr	Gly	Ala	
	Thr		115					120				*	125	_		
35	,	*					•									
	Leu															
40	<210 <400		.37 < .37	(211>	· 41	.5 <2	12>	PRT	<21	.3>	Esch	eric	chia	coli		
	Met Thr	Lys	Arg	Lys	His	Leu	Leu	Leu	Leu	Leu	Leu	Phe	Ser	Phe	Ser	

45 1 5 5 10 15

Asn Ser Ala Pro Leu Tyr Ser Leu Ile Arg Glu Ala Val Met His
Asp
50 20 25 30

315/370

Pro Ile Val Met Glu Ala Arg Ala Glu Leu Thr Ser Ala Gln Ser Arq Ile Glu Gln Ala Ser Ser Ala His Trp Pro Val Val Thr Ala Thr Gly Ser Lys Leu Leu Ser Gln Ser His Arg Tyr Ser Tyr Asp Tyr Asp Thr Glu Asp Ile Leu Pro Gly Ile Arg Gly Glu Val Asn Ile Phe Ala Ser Gly Ala Ile Glu Ala Asp Val Arg Arg Ser Glu Ser Glu Ala Glu Tyr His Tyr Lys Met Glu Glu Thr Lys Glu Glu Thr Ile His Ser Phe Val Ser Leu Tyr Leu Asp Ala Leu Arg Glu Lys Gln Ser Ile Ala Val Leu Glu Gln Ser Leu Ser Arg His Asn Ala Ile Leu Asn Asp Leu Asn Thr Ile Ser Ile His Asp Thr Gly Arq Glu Ser Glu Leu Val Gln Ala Glu Ala Arg Arg Leu Met Val Arg Gln Gln Ile Asn Ser Arg Ser Arq

								310/.	370						
	Val Pro	Leu	Lys	Thr	Thr	Leu	Gly	Lys	Leu	Ser	Thr	Trp	Thr	Lys	Asn
5			195					200					205		
	Val Ala	Thr	Glu	Ala	Asp	Leu	Glu	Asn ·	Pro	Phe	Ser	Arg	Met	Thr	Glu
10		210					215					220			
	Lys Ser	Leu	Leu	Thr	Asp	Phe	Thr	Gln	Ala	Pro	Gln	Lys	Gly	Asn	Pro
15	225 240					230		¥)			235				
	Trp Lys	Leu	Ala	Ser	Gln	Ala	Asp	Val	Glu	Ser	Lys	Lys	Ala	Ala	Leu
20	175				245					250					255
	Ala Val	Gln	Glu	Leu	Ala	Arg	Tyr	Pro	Arg	Val	Asp	Leu	Thr	Gly	Ser
25				260			-		265					270	
	Thr Phe	Arg	Asp	Asp	Gln	Gln	Ile	Gly	Val	Asn	Leu	Ser	Trp	Asp	Leu
30			275					280	•		١		285		
	Asn Val	Arg	Asn	Ala	Ser	Tyr	Gly	Val	Thr	Glu	Lys	Ala	Ala	Gln	Ile
35		290					295		· -			300			
	Ala Thr	Ala	Thr	Gly	Arg	Leu	Asp	Ser	Val	Ala	Arg	Met	Ile	Asp	Glu
40	305 320					310					315				
45	_	Arg	Leu	Ser	Leu	Ile	Thr	Val	Arg	Gln	Ser	Arg	Gly	Glu	Met
73	Glu				325					330					335

Thr Leu Arg Arg Gln Glu Gln Ala Ser Ala Arg Val Val Asp Phe 50 Tyr 340 Ser 345 350

								,								
	Arg		Gln	Phe	Gln	Val	Ala	Arg	Lys	Thr	Leu	Ile	Glu	Leu	Leu	
5	11011		355					360				,	365			٠
	Ala	Glu	Asn	Glu	Leu	Tyr	Ser	Val	Gly	Leu	Ser	Arq	Val	Gln	Thr	
10	Glu	370				_	375	-	-			380				
						-	-				-				•	
	Asp Met	Gln	Met	Leu	His	Gly	Met	Leu	Asp	Tyr	Leu	Tyr	Ser	Gln	Gly	
15	385 400					390		•			395				ı	
20	Leu 	Leu	Lys	Trp	Ser 405	Gly	Val	Asn	Leu	Ser 410	Gly	Glu	Glu	Glu	Lys 415	
•	<210) \	130 /	~911\		71 /	2125	PR:	n /ð-	1 2 5	T = -1		1*			
25	<400		L38	· Z I I .) <u>.</u> \ /		FR.	L \Z.	13/	ESCI	ieric	cnia	coli	- . ·	
23	Met Ser	Lys	Phe	Leu	Pro	Leu	Leu	Ala	Leu	Leu	Ile	Ser	Pro	Phe	Val	. :
	1				5					10					15 '	
30	 Ala	T.e.ii	Thr	T,e11	Asn	Asn	T.e.:	Gl n	Gln	Δκα	Phe	Thr			Dwo	
	Val	дса	J.117	20	7150	мэр		GIII	25	Arg	rne	, TIIT	GIU,		· ·	
35				20					23					30		
· .	Ile Gln	Arg	Ala	His	Phe	Asp	Gln	Thr	Arg	Thr	Ile	Lys	Asp	Leu	Pro	
	OIII		35					40					45			
40	Pro	T 011	7\ ~~	C0.2	Cln	C1	Cl~	Ma +	T 0	T7.	ת ד ת	,		0 1		
	Leu		Arg	ser	GTII	дтХ		Met	цец	тте	АТА		Asp	GIn	· GTÀ	
45		50					55 .					60				
-r <i>J</i>		Trp	Asp	Gln	Thr	Ser	Pro	Phe	Pro	Met	Gln	Leu	Leu	Leu	Asp	
	Asp 65			•		70					75					80

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	V	WO 200	05/0978	323				318	PCT/EP2005/00397						
	Lys Thr	Arg	Met	Val	Gln 85	Val	Ile	Asn	Gly	Gln 90	Pro	Pro	Gln	Ile	
5	· Zla	Glu	Z) en	Asn		Gln	Mot	Pho	Cln		7) a n	II	Т о		95
	Ala		ASII	100	110	GIII	mec	riie	105	rne	Asn	urs	теп	ьеи 110	Arg
10	Leu Glu	·Phe	Gln	Ala	Asp	Arg	Lys	Val	Leu	Glu	Gln	Asn	Phe	Arg	Val
	Oru		115					120		,			125	,	,
15	Phe Thr		Asp	Lys	Gly	Gl _. u	Gly	Arg	Trp	Thr	Leu	Arg	Leu	Thr	Pro
20	·	130					135					140			
	Lys	Ţhr	Pro	Leu	Asp	•	Ile	Phe	Asn	Thr	Ile	Asp	Leu	Ala	Gly
25	145 160					150				1	155				•
	Thr Thr	Tyr	Leu	Glu	Ser	Ile	Gln	Leu'	Asn	Asp	Lys,	Gln	Gly	Asp	Arg
30					165				÷	170					175
	Asp Asp	Ile	Ala	Leu 180	Thŗ	Gln	His	Gln	Leu 185	Thr	Pro	Ala	Gln	Leu 190	Thr
35	_													190	
	Asp	GLu	H1S 195	GIn	Arg	Phe	Ala	Ala 200	Gln						
40	-01		20.	·011\	. 75		1105	22.		2.			, ,		

139 <211> 770 <212> PRT <213> Escherichia coli <210> <400> Met Glu Asn Phe Phe Met Lys Asn Ser Lys Val Phe Tyr Arg Ser

Ala

Leu Ala Thr Ala Ile Val Met Ala Leu Ser $extst{Ala}$ Pro Ala Phe Ala . Thr

		Ser	Thr	Val	Ser	Thr	Asp	Pro	Val	Thr	Leu	Asn	Thr	Glu	Lys	· ·
5	Thr		35 .		·			40					45	٠		
	Thr Ala	Leụ	Asp	Gln	Asp	Val	Val	Ile	Asn	Gly	Asp	Asn	Lys	Ile	Thr	
10		50					55					60				
	Val Phe	Thr	Ile	Glu	Thr	Ser	Asp	Ser	Asp	Lys	Asp	Leu	Asn	Val	Thr	
15	65	,		,		70					75			•		80
	Gly Val	Gly	His	Asp	Ile	Thr	Ala	Ala	Ser	Thr	Val	Asn	Gln	Asp	Phe	
20	• • • • • • • • • • • • • • • • • • • •				85			٠.		90					95	
,	Glu Thr	Gly	Val	Lys	Val	Ser	Gly	Asn	Lys	Asn	Val	Val	Ile	Asn	Ala	
25				100					105					110		
	Asp Ala	Ser	Thr	Ile	Thr _.	Ala	Gln	Gly	Glu	Gly	Thr	T yr	Val	Arg	Thr	
30	211.U		115					120					125	04		
	Met Phe	Val	Ile	Asp	Ser	Thr	Gly	Asp	Val	Val	Val	Asn	Gly	Gly	Asn	
35	1110	130		•			135			*		140				
	Val Ala	Ala	Lys	Asn	Glu	Lys	Glý	Ser	Ala	Thr	Gly	Ile	Ser	Leu	Glu	
40	145 160					150					155		•			
	Thr.	Thr	Cl v	7) an	Nen	T.O.	ሞኮኮ	T 011	, Nan	e. Gly	Thr.	Thr	Tlo	7 an	7\ 1 ¬	
45	Gln	1117	Gry	ASII	165	,		nea	VPII	170	T11T	1 1111	116	ASII	175	-
							,									
50	Gly Lys	Asn	Lys	Ser 180	Tyr	Ser	Asn	Gly	Ser 185	Thr	Ala	Ile	Phe	Ala 190	Gln	
				100					100					4,00		

	Gly Thr	Asn	Leu	Leu	Gln	Gly	Phe	Asp	Gly	Asp	Ala	Thr	Asp	Asn	Ile
5			195			. •		200			,		205		
	Leu Thr		Asp	Ser	Asn	Ile	Ile	Asn	Gly	Gly	Ile	Glu	Thr	Ile	Val
10		210					215			,		220			
15	Ala Asp	Gly	Asn	Lys	Thr	Gly	Ile	His	Thr	Val	Asn	Leu	Asn	Ile	Lys
	225					230					235				
20	Gly Ser	Ser	Val [.]	Ile	Gly	Ala	Ala	Asn	Asn	Lys	Gln	Thr	Ile	Tyr	Ala
			•		245					250					255
	Ala Ser	Ser	Ala	Gln	Gly	Ala	Gly	Ser	Ala	Thr	Gln	Asn	Leu	Asm	Leu
				260			•		265					27 O	
30	Val Ser	Ala	Asp	Ser	Thr	Ile	Tyr	Ser	Asp	Val	Leu	Ala	Leu	Sex	Glu
			275	•			2	280					285		
35	Glu Arg	Asn	Ser	Ala	Ser	Thr	Thr	Thr	Asn	Val	Asn	Met	Asn	Vаl	Ala
	9	290					295	,				300			
40	Ser Ala	Tyr	Trp	Glu	Gly	Asn	Ala	Tyr	Thr	Phe	Asn	Ser	Gly	Asp	Lys
	305 320					310					315				
45	Gly Gly	Ser	Asp	Leu	Asp	Ile	Asn	Leu	Ser	Asp	Ser	Ser.	Val	Trp	Lys
	-				325			•		330					335
50	Lys	Val	Ser	Gly	Ala	Gly	Asp	Ala	Ser	Val	Ser	Leu	Gln	Asn	Gly

Ser

340 345 350

Val Trp Asn Val Thr Gly Ser Ser Thr Val Asp Ala Leu Ala Val 5 Lys

355 360 365

Asp Ser Thr Val Asn Ile Thr Lys Ala Thr Val Asn Thr Gly Thr 10 Phe 370 375 380

Ala Ser Gln Asn Gly Thr Leu Ile Val Asp Ala Ser Ser Glu Asn 15 Thr

385 390 395 400

420

20 Leu Asp Ile Ser Gly Lys Ala Ser Gly Asp Leu Arg Val Tyr Ser Ala
405 410 415

25 Gly Ser Leu Asp Leu Ile Asn Glu Gln Thr Ala Phe Ile Ser Thr Gly

425

430

30 Lys Asp Ser Thr Leu Lys Ala Thr Gly Thr Thr Glu Gly Gly Leu

Tyr
435
440
445

35 Gln Tyr Asp Leu Thr Gln Gly Ala Asp Gly Asn Phe Tyr Phe Val Lys 450 455 460

40 Asn Thr His Lys Ala Ser Asn Ala Ser Ser Val Ile Gln Ala Met Ala 465 470 475

45
Ala Ala Pro Ala Asn Val Ala Asn Leu Gln Ala Asp Thr Leu Ser Ala
485
490
495

322/37

	Arg Trp	Gln	Asp	Ala	Val	Arg	Leu	Ser	.Glu	Asn	Asp	Lys	Gly	Gly	Val
		•		500					505					510	
5 `		Gln			Gly	Gly	Lys		Lys	His	Thr	Thr		Gly	Asn
10		·	515	•				520					525		
10	Ser Arg	Tyr	Asp	Leu	Asp	Val	Asn	Gly	Val	Met	Leu	Gly	Gly	Asp	Thr
		530					535					540			
15	Phe	Met	Thr	Glu	Asp	Gly	Ser	Trp	Leu	Ala	Gly	Val	Ala	Met	Ser
	Ser 545 560					550			,		555				
20	*														*
	Ala Gly	Lys	Gly	Asp	Met	Thr	Thr	Met	Gln	Ser	Lys	Gly	Asp	Thr	Glu
25					565					570				•	575
	Tyr Phe	Ser	Phe	His	Ala	Tyr	Leu	Ser	Arg	Gln	Tyr	Asn	Asn	Gly	Ile
30				580					585				٠	590	
	Ile Val	Asp	Thr	Ala	Ala	Gln	Phe	Gly	His	Tyr	Ser	Asn	Thr	Ala	Asp
35			595					600					605		
	Arg Asn	Leu	Met	Asn	Gly	Gly	Gly	Thr	Ile	Lys	Ala	Asp	Phe	Asn	Thr
40		610					615					620			
	Gly Asn	Phe	Gly	Ala	Met	Val	Lys	Gly	Gly	Tyr	Thr	Trp	Lys	Asp	Gly
45	625 640					630					635				
	Gly	Leu	Phe	Ile	Gln	Pro	Tyr	Ala	Lys	Leu	Ser	Ala	Leu	Thr	Leu
50	Glu				645					650				÷	655

			•												
		Val	Asp	Tyr	Gln	Leu	Asn	Gly	Val	Asp	Val	His	Ser	Asp	Ser
5	Tyr _.			660					665					670	٠.
111	Asn Ala	Ser	Val	Leu	Gly	Glu	Ala	Gly	Thr	Arg	Val	Gly	Tyr	Asp	Phe
10	1414		675					680					685		
		Gly	Asn	Ala	Thr	Val	Lys	Pro	Tyr	Leu	Asn	Leu	Ala	Ala	Leu
15	Asn	690					695					700		ı	
	Glu Asn	Phe	Ser	Asp	Gly	Asn	Lys	Val	Arg	Leu	Gly	Asp	Glu	Ser	Val
20	705 720					710					715				
		Ser	Ile	Asp	Gly	Ala	Ala	Phe	Arg	Val	Gly	Ala	Gly	Val	Gln
25	Ala				725					730					735
		Ile	Thr	Lys	Asn	Met	Gly	Ala	Tyr	Ala	Ser	Leu	Asp	Tyr	Thr
30	Lys		,	740					745					750	
	Gly Val	Asp	Asp	Ile	Glu	Asn	Pro	Leu	Gln	Gly	Val	Val	Gly	Ile	Asn
35	vaı		755					760					765		
40	Thr	Trp 770				J									
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15	N/1 1-	0 -	7	D	~ 3	D1	m1.	0 -	•	71	τ -	Q -		T	T. T

45 Met Ser Arg Pro Gln Phe Thr Ser Leu Arg Leu Ser Leu Leu Ala Leu
1 5 10 15

50 Ala Val Ser Ala Thr Leu Pro Thr Phe Ala Phe Ala Thr Glu Thr Met

20 25 30

Thr Val Thr Ala Thr Gly Asn Ala Arg Ser Ser Phe Glu Ala Pro 5 Met 40 45

Met Val Ser Val Ile Asp Thr Ser Ala Pro Glu Asn Gln Thr Ala

10 Thr
50 55 60

Ser Ala Thr Asp Leu Leu Arg His Val Pro Gly Ile Thr Leu Asp
15 Gly
65 70 75 80

Thr Gly Arg Thr Asn Gly Gln Asp Val Asn Met Arg Gly Tyr Asp 20 His 85 90 95

Arg Gly Val Leu Val Leu Val Asp Gly Val Arg Gln Gly Thr Asp
25 Thr
100 105 110

Gly His Leu Asn Gly Thr Phe Leu Asp Pro Ala Leu Ile Lys Arg

Val

115

120

125

Glu Ile Val Arg Gly Pro Ser Ala Leu Leu Tyr Gly Ser Gly Ala 35 Leu 130 . 135 140

45 Glu Gly Gln Ser Ser Gly Phe Arg Val Phe Gly Thr Gly Gly Thr Gly
165 170 • 175

 $50~{\rm Asp}$ His Ser Leu Gly Leu Gly Ala Ser Ala Phe Gly Arg Thr Glu Asn

325/370

				180					185	•				190	
- 5	Leu Arg	Asp	Gly 195	Ile	Val	Ala	Trp	Ser 200	Ser	Arg	Asp	Arg	Gly 205	Asp	Leu
10	Gln Met	Ser 210	Asn	Gly	Glu	Thr	Ala 215	Pro	Asn	Asp	Glu	Ser 220	Ile	Asn	Asn
15	Leu Gly 225 240	Ala	Lys	Gly	Thr	Trp 230	Gln	Ile	Asp	Ser	Ala 235	Gln	Ser	Leu	Ser
20	Leu Gln	Val	Arg	Tyr	Tyr 245	Asn	Asn	Asp	Ala	Arg 250	Glu	Pro	Lys	Asn	Pro [.] 255
25	Thr Thr	Val	Glu	Ala 260	Ser	Asp	Ser	Ser	Asn 265	Pro	Met	Val	Asp	Arg 270	Ser
30	Ile Asn	Gln	Arg 275	Asp	Ala	Gln	Leu	Ser 280	Tyr	Lys	Leu	Ala	Pro 285	Gln	Gly
35	Asp Ile	Trp	Leu	Asn	Ala	Asp	Ala 295	Lys	Ile	Tyr	Trp	Ser 300	Glu	Val	Arg
40	Asn Thr 305 320	Ala	Gln	Asn	Thr	Gly 310	Ser	Ser	Gly	Glu	Tyr 315	Arg	Glu	Gln	Ile
45	Lys Phe	Gly	Ala	Arg	Leu 325	Glu	Asn	Arg	Ser	Thr 330	Leu	Phe	Ala	Asp	Ser 335

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	W	O 200	5/09782	23				20 (/2)	70				PCT/I	E P2 005	5/003972
								326/3							
	Ala Gln	Ser	His		Leµ	Thr	Tyr	Gly	_	Glu	Tyr	Tyr	Arg	Gln	Glu
				340					345					350.	. 4
5			Gly	Gly	Ala	Thr	Thr	Gly	Phe	Pro	Gln	Ala	Lys	Ile	Asp
	Phe		355					360					365		
10	C	C	C1	M 1010	Ton	Cl n	7 ~~	Clas	710	mh m	Т олг	71 20 00	7	т	Dead
	Ile	370	Gly	Trp	теп.	GIII	375	GIU	тте	THE	теп	380	Asp	ьец	Pro.
15		370			1		373					300			
13	Thr Asp	Leu	Leu	Gly	Gly	Thr	Arg	Tyr	Asp	Ser	Tyr	Arg	Gly	Ser	Ser
	385 400					390					395				
20				*	•								•		
	Gly Met	Tyr	Lys	Asp	Val	Asp	Ala	Asp	Lys	Trp	Ser	Ser	Arg	Ala	Gly
25					405					410					415
	Thr Gln	Ile	Asn	Pro	Thr	Asn	Trp	Leu	Met	Leu	Phe	Gly	Ser	Tyr	Ala
30	0111			420					425	*				430	
	Ala His	Phe	Arg	Ala	Pro	Thr	Met		Glu	Met	Tyr	Asn		Ser	Lys
35			435					440					445		
		Ser	Ile	Gly	Arg	Phe	Tyr	Thr	Asn	Tyr	Trp	Val	Pro	Asn	Pro,
40	Asn	450					455			•		460		٠,	
40			•												
	Leu Arg	Arg	Pro	Glu	Thr	Asn	Glu	Thr	Gln	Glu	Tyr	Gly	Phe	Gly	Leu
45	465					470					475	٠			
	Phe	Asp	Asp	Leu	Met	Leu	Ser	Asn	Asp	Ala	Leu	Glu	Phe	Lys	Ala
50	Ser	*	, -		485				_	490					495
															•

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Tyr Phe Asp Thr Lys Ala Lys Asp Tyr Ile Ser Thr Thr Val Asp Phe Ala Ala Ala Thr Thr Met Ser Tyr Asn Val Pro Asn Ala Lys Ile Trp Gly Trp Asp Val Met Thr Lys Tyr Thr Thr Asp Leu Phe Ser Leu Asp Val Ala Tyr Asn Arg Thr Arg Gly Lys Asp Thr Asp Thr Gly Glu Tyr Ile Ser Ser Ile Asn Pro Asp Thr Val Thr Ser Thr Leu Asn Ile Pro Ile Ala His Ser Gly Phe Ser Val Gly Trp Val Gly Thr Phe Ala Asp Arg Ser Thr His Ile Ser Ser Ser Tyr Ser Lys Gln Pro Gly Tyr Gly Val Asn Asp Phe Tyr Val Ser Tyr Gln Gly Gln Gln Ala Leu Lys Gly Met Thr Thr Leu Val Leu Gly Asn Ala Phe Asp Lys Glu Tyr Trp Ser Pro Gln Gly Ile Pro Gln Asp Gly Arg Asn Gly Lys Ile Phe Val

Ser Tyr Gln Trp 660

5

<210> 141 <211> 719 <212> PRT <213> Escherichia coli <400> 141

- 10 Met Arg Asp Glu Met Leu Tyr Asn Ile Pro Cys Arg Ile Tyr Ile Leu 1 5 10 15
- 15 Ser Thr Leu Ser Leu Cys Ile Ser Gly Ile Val Ser Thr Ala Thr Ala
 20 25 30
- 20 Thr Ser Ser Glu Thr Lys Ile Ser Asn Glu Glu Thr Leu Val Val Thr 35 40 45
- 25 Thr Asn Arg Ser Ala Ser Asn Leu Trp Glu Ser Pro Ala Thr Ile Gln 50 55 60
- 30 Val Ile Asp Gln Gln Thr Leu Gln Asn Ser Thr Asn Ala Ser Ile Ala 65 70 75 80
- 35 Asp Asn Leu Gln Asp Ile Pro Gly Val Glu Ile Thr Asp Asn Ser Leu
 85 90 95
- 40 Ala Gly Arg Lys Gln Ile Arg Ile Arg Gly Glu Ala Ser Ser Arg Val 100 105 110
- 45 Leu Ile Leu Ile Asp Gly Gln Glu Val Thr Tyr Gln Arg Ala Gly Asp
 115 120 125
- 50 Asn Tyr Gly Val Gly Leu Leu Ile Asp Glu Ser Ala Leu Glu Arg Val

130 135 140

- Glu Val Val Lys Gly Pro Tyr Ser Val Leu Tyr Gly Ser Gln Ala

 5 Ile
 145 150 155
 160
- Gly Gly Ile Val Asn Phe Ile Thr Lys Lys Gly Gly Asp Lys Leu Ala
 165 170 175
- Ser Gly Val Val Lys Ala Val Tyr Asn Ser Ala Thr Ala Gly Trp Glu
 180 185 190
- 20 Glu Ser Ile Ala Val Gln Gly Ser Ile Gly Gly Phe Asp Tyr Arg Ile 195 200 205
- 25 Asn Gly Ser Tyr Ser Asp Gln Gly Asn Arg Asp Thr Pro Asp Gly Arg . 210 . 215 . 220
- Leu Pro Asn Thr Asn Tyr Arg Asn Asn Ser Gln Gly Val Trp Leu Gly 225 230 235
- Tyr Asn Ser Gly Asn His Arg Phe Gly Leu Ser Leu Asp Arg Tyr Arg

 245
 250
 255
- 40
 Leu Ala Thr Gln Thr Tyr Tyr Glu Asp Pro Asp Gly Ser Tyr Glu Ala
 260
 270
- Phe Ser Val Lys Ile Pro Lys Leu Glu Arg Glu Lys Val Gly Val Phe

 275 280 285

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Tyr Asp Thr Asp Val Asp Gly Asp Tyr Leu Lys Lys Ile His Phe Asp Ala Tyr Glu Gln Thr Ile Gln Arg Gln Phe Ala Asn Glu Val Lys Thr Thr Gln Pro Val Pro Ser Pro Met Ile Gln Ala Leu Thr Val His Asn Lys Thr Asp Thr His Asp Lys Gln Tyr Thr Gln Ala Val Thr Leu Gln Ser His Phe Ser Leu Pro Ala Asn Asn Glu Leu Val Thr Gly Ala Gln Tyr Lys Gln Asp Arg Val Ser Gln Arg Ser Gly Gly Met Thr Ser Ser Lys Ser Leu Thr Gly Phe Ile Asn Lys Glu Thr Arg Thr Arg Ser Tyr 400. Tyr Glu Ser Glu Gln Ser Thr Val Ser Leu Phe Ala Gln Asn Asp Trp Arg Phe Ala Asp His Trp Thr Trp Thr Met Gly Val Arg Gln Tyr Trp 420-Leu Ser Ser Lys Leu Thr Arg Gly Asp Gly Val Ser Tyr Thr Ala Gly

	,														
5	Ile Glu	Ile 450		: Asp	Thr	: Ser	Leu 455		a Arç	g Glu	ı Ser	Ala 460		Asp	His
10	Met Glu 465 480		. Thr	Ser	Thr	Ser 470		Arg	J Tyr	: Ser	Gly 475	Phe	Asp	Asn	Leu
10	Leu	Arg	Ala	Ala	Phe	Ala	Gln	Gly	Tyr	· Val	Phe	Pro	Ťhr	Leu	Ser
15	Gln				485					490					495
20	Leu Pro	Phe	Met	Gln 500	Thr	Ser	Ala	Gly	Gly	•	Val	Thr	Tyr	Gly 510	Asn
25	Asp Tyr	Leu	·Lys 515	Ala	Glu	His	Ser	Asn 520	Asn		Glu	Leu	Gly 525		Arg
	Asn Ala			Thr	Trp	Leu	Ile			Ala	Val	Tyr		Ser	Glu
. 30	Ture	530	777 Y	T 1 0	7. 7. 7.	Com	535	T]_		70	•	540			
35	Asn 545 560		·		AIA	Ser 550	ьеи	TTE	Cys	Asp	.GLY	Ser	Ile	Val	Cys.
40	Gly Ile	Asn	Thr	Asņ	Ser 565	Ser	Arg	Ser	Ser	Tyr 570	Tyr	Tyr	Tyr		Asn 575
45	Ásp Gly	Arg	Ala	Lys 580	Thr	Trp	Gly	Leu	Glu 585	Ile	Ser	Ala		Ту <u>г</u> 590	Asn
50	Trp Tyr	Val	Phe 595	Ser	Pro	Tyr	Ile	Ser	Gly	Asn	Leu	Ile	Arg 605	Arg	Gln

	Glu Asn	Thr	Ser	Thr	: Leu	Lys	Th	r Thr	Asn	Thr	Gly	Glu	Pro	Ala	Ile
. 5		610					615	5				620			
	Glv	Ara	Tle	Glv	т Т.еп	Tvvs	. His	s Thr	T. 611	77 ~ 7	Mot	C1	C 7	7N T _	70
10	Ile 625			1		630			шси	. vai	635	GTĀ	GTII	Ara	Asn
•	640					,					033				
í	Ile	Ser	Asp	Val	Phe	Tle	Arc	g Ala	7\ 1 ->	802	Con	70 7 ~	T	70	77.
15	Ser				645		111.0		ALA	650	per	Ald	пуѕ	Asp	
					010					630					655
20	Asn Ala	Gly	Thr	Glu	Thr	Asn	Val	. Pro	Gly	Trp	Ala	Thr	Leù	Asn	Phe
	1120			660					665		*			670	
	Val	Asn	Thr	G] 11	Pha	Glu	λεν	Glu	7	C1 ~	C	71	~ 7		
25	Ala		675	Olu	1110	Ory	ASII	680	ASP	GTII	ser	Arg		Asn	Leu
	•		0,0					000					685		
30	Leu Ile	Asn	Asn	Leu	Thr	Asp	Lys	Arg	Tyr	Arg	Thr	Ala	His	Glu	Thr
		690					695				•	700			
•	Pro	Δla	Δla	Glv	Phe	Nan	ת ד ת	7\]_	T] _	C1 '	D1	77 J	-	_	
35	705	111.C	1114	O _T y	1116	710	АТА	<i>A</i> la	тте	GТĀ	715	vaı	Trp	Asn	Phe
	<210	_ 1	12 /	·011'\	. 10	10 70	010	DD		2.	· 1				
40	<400	> 1	.42	. 4. 4. 4. 7	7 13	79 \2	.14/	PRI	,21	.3>	Escn	eric.	nıa	coli	· · · · · · · · · · · · · · · · · · ·
10	Met . Ser	Arg	Lys	Val	Cys	Ala	Val	Ile	Leu	Ser	Ala	Ala	Ile	Cys	Leụ
	1				5					10					15
45	17 a 1	2 0∞	C1	71 7	D == =	71 7 -	(TI) = - : -	70.7	a	0 3	1	. .			
	Ser	ser	σтλ	Ата	rro	нта	Trp	Ala	Ser	GLu	His	Gln	Ser	Thr	Leu

. 25

30

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•	333/3/0	

	Ala Leu	Gly	Tyr	Leu	His	Ala	Arg	Thr	Asn	Ala	Pro	Gly	Ser	Asp	Asn	
	пси		35					40					45			,
5 ·	Asn	Glv	Tle	Asn	Val	Tivs	Tyr	Arg	Tyr	Glu	Phe	Thr	Asn	Δla	T.611	
	Gly	50			, , ,	_, _	55	¥ 9	-1-	014	1110	60	пор	111.0	шси	
10				*												
	Leu Thr	Ile	Thr	Ser	Phe	Ser	Tyr	Ala	Asn	Ala	Glu	Asp	Glu	Gln	Lys	
	65		•			70					75					80
15	His	Tyr	Ser	Asp	Thr	Arg	Trp	His	Glu	Asp	Ser	Val	Arg	Asn	Arq	
	Trp				85	_				90					95	
20						•										
•	Phe Ser	Ser	Val	Met	Ala	Gly	Pro	Ser	Val	Arg	Val	Asn	Glu	Trp	Phe	
	8		-	100					105					110		
25		Tyr	Ser	Met	Ala	Gly	Val	Ala	Tyr	Ser	Arg	Val	Ser	Thr	Phe	
·	Ser		115					120			,		125			
30										•						
	Gly Val		T _. yr	Leu	Arg	Val		Asp	Asn	Lys	Gly		Thr	His	Asp	
0.5		130					135				·	140	×			
35		Thr	Gly	Ser	Asp	Asp	Gly	Arg	His	Ser	Asn	Thr	Ser	Leu	Ala	
	Trp 145					150					155					
40	160					•										
		Ala	Gly	Val	Gln	Phe	Asn	Pro	Thr	Glu	Ser	Val	Thr	Ile	Asp	
15	Leu				165					170			•		175	
45			~ 1	~ 7	~			.	_			_,				•
	Ala Ile	туг	GLu		ser	GTĀ	Ser	Gly		Trp.	Aŗg	Thr	Asp		Phe	
50				180					185					190		

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Val Gly Ile Gly Tyr Arg Phe 195

5 <210> 143 <211> 456 <212> PRT <213> Escherichia coli <400> 143

Met Lys Lys Ser Thr Leu Ser Leu Ala Ile Gly Leu Leu Leu Ala Cys
10 1 5 10 15

Ser Thr Gly Met Ala Lys Thr Gln His Leu Thr Leu Glu Gln Arg Leu - 25 30

Glu Ala Ala Glu Met Arg Ala Ala Lys Ala Glu Gly Gln Val Lys Gln 20 35 40 45

Leu Gln Thr Gln Gln Ala Ala Glu Ile Arg Glu Ile Lys Thr Ala Gln 25 50 55 60

Gly Asn Thr Pro Val Asn Gly Gln Ser Thr Thr Glu Ser Glu Lys Lys 30 65 70 75 80

Asn Ala Thr Pro Pro Asn Leu Leu Leu Ser Gly Tyr Gly Asp Leu Lys
35 85 90 95

Ile Tyr Gly Asp Val Glu Phe Asn Met Asp Ala Glu Ser Asn His Gly
40 100 105 110

Leu Leu Ala Met Thr Asn Ala Asp Val Asn Ser Asp Pro Thr Asn Glu
45 ' 115 120 125

Trp Asn Leu Asn Gly Arg Ile Leu Leu Gly Phe Asp Gly Met Arg
Lys
50 130 135 140

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WO 200	5/097823		33	5/370				РСТ	/EP200	5/003972
Leu Asp Asp 145 5 160	Asn Gly	Tyr Phe		y Phe		Ala 155	Gln	Pro	Leu	Gly
Met His Lys	Gly Ser	Val Asn	Ile As	p Asp	Ala 170	Val	Phe	Phe	Phe	Gly _. 175
Glu Asn Met 5	Asp Trp	Lys Val	Lys Va	l Gly 185	Arg	Phe	Glu	Ala	Tyr 190	Asp
Phe Pro Ala	Leu Asn	Gln Asp	Thr Phe		Glu [°]	His	Ser	Gly 205	Asn	Thr
Asn Asp Glu 5 210	Leu Tyr	Asp Asp	Gly Ser	c Gly	Tyr	Ile	Tyr 220	Met	Met	Lys
Gly Arg Gln 0 225 240	Gly Arg	Ser Asn 230	Ala Gly	g Gly	Asn	Phe 235	Lėu	Val	Ser	Lys
Leu Asp Thr	Asn Trp	Tyr Phe	Glu Lei	ı Asn	Thr 250	Leu	Leu	Glu	Asp	Gly 255
Ser Leu Gln	Tyr Asn 260	Asp Gly	Asn Tyr	265		Arg	Asp	Met	Glu 270	Gln
Lys Asn 5 Glu	Val Ala 275	Tyr Leu	Arg Pro		Ile	Ala	Trp	Ser 285	Pro	Thr
Glu Phe Ala . 290	Thr Val	Ser Ala	Ala Met	Glu	Ala	Asn	Val	Val	Asn	Asn

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Tyr Gly Tyr Thr Asp Ser Lys Gly Asn Phe Val Asp Gln Ser Asp 310 -Thr Gly Tyr Gly Met Ser Met Thr Trp Asn Gly Leu Lys Thr Asp Glu Asn Gly Ile Val Val Asn Leu Asn Thr Ala Tyr Leu Asp Ala Asn 35 0 Asn Glu Lys Asp Phe Thr Ala Gly Ile Asn Ala Leu Trp Lys Arg Phe Glu Leu Gly Tyr Ile Tyr Ala His Asn Lys Ile Asp Glu Phe Ser Gly Val Val Cys Asp Asn Asp Cys Trp Ile Asp Asp Glu Gly Thr Tyr Asn Ile His Thr Ile His Ala Ser Tyr Gln Phe Ala Asn Val Met Asp Met Glu Asn Phe Asn Ile Tyr Leu Gly Thr Tyr Tyr Ser Ile Leu Asp Ser 43 O Asp Gly Asp Lys Ile His Gly Asp Asp Ser Asp Asp Arg Tyr Gly Ala Arg Val Arg Phe Lys Tyr Phe Phe

5	<21 <40		144 144	<211	> 1	74 <	212>	PR	T <2	13>	Esc	heri	chia	col.	i	
3			Gly	Lys	Ala	Phe	Leu	Ala	Cys	Val	Leu	Met	Ser	Val	Val	
	Leu 1				5					10					15	
10																
	Thr Pro	Gly	Cys	Glu	Thr	Ala	Lys	Lys	Ile	Ser	Gln	Val	Ile	Arg	Asn	
				20					25					30 .		
15	Asp	Ile	Gln	Val	Gly	Lvs	Leu	Met	Asp	Gln	Ser	Thr	Glu	T.e.11	Thr	
	Val		35		1	-1-		40			201	2111	45	Lea	1111	
20			33		÷			4,0	٠				40			
20		Leu	Leu	Thr	Glu	Pro	Asp	Ser	Asn	Leu	Thr	Ala	Asp	Gly	Glu	
	Ala	50					55					60			•	
25																
	Ala Phe	Pro	Val	Asp	Val	Gln	Leu	Val	Tyr	Leu	Ser	Asp	Asp	Ser	Lys	
	65					70					75					80
30	His	Ala	Ala	Asp	Tyr	Asp	Gln	Val	Ala	Thr	Thr	Ala	Leu	Pro	Asp	
	Val			_	85	-				90					95	٠
35																
50		Gly	Lys	Asn	Tyr	Ile	Asp	His	Gln	Asp	Phe	Asn	Leu	Leu	Pro	•
	Asp			100					105				•	110		
40	em 1		_			_	_		_	_	_					
	Thr Tyr	Val		Thr	Leu	Pro	Pro		Lys	Leu	Asp	Glu	Lys	Thr	Gly	
			115					120					125			
45	Ile	Gly	Val	Ile	Ala	Tyr	Phe	Ser	Asp	Asp	Gln	Ala	Thr	Glu	Trp	
	Lys	130					135		-	-		140			_	
50								•								
										•						

	W	/ O 2 00	5/09782	23				338/3	370				PCT	/EP200	05/0039	72
5	Gln Val 145 160	Ile	Glu	Ser	Val	Glu 150	Ser	Ile	Gly	His	His 155	Tyr	Arg	Leu	Leu	
	His	Iľe	Arg	Ala	Ser 165	Ala	Ile	Glu	Met	Lys 170	Lys	Glu	Glu	Asn		
10	<210 <400		145 < 145	<211>	> 11	L44 ,<	<212>	> PI	RT <2	213>	Esc	cheri	ichi	a col	Li	
15	Leu Gly 1	Thr	Leu	Ala	Trp 5	Ile	Phe	Leu	Leu	Val	Trp	Ile	Trp	Trp	Gln 15	
. 20	Pro Asn	Lys	Trp	Thr	Leu	Tyr	Glu	Gln	His	Trp	Leu	Ala	Pro	Leu 30	Ala	
25	Arg Leu	Trp	Leu 35	Ala	Thr	Ala	Val	Trp	Gly	Leu	Ile	Ala	Leu 45	Val	Trp	
30	Thr Lys	Trp	Arg	Val	Met	Lys	Arg 55	Leu	Gln	Lys	Leu ,	Glu 60	Lys	Gln	Gln	
35	Gln Arg 65	Gln	Arg	Glu	Glu	Glu 70	Lys	Asp	Pro	Leu	Thr	Val	Ģlu	Leu	His	80
40	Gln Leu	Gln	Glņ	Tyr	Leu 85	Asp	His	Trp	Leu	Leu 90	Arg	Leu	Arg	Arg	His 95	
45	Asp Gly	Asn	Arg	Arg	Tyr	Leu	Trp	Gln	Leu 105	Pro	Trp	Tyr	Met	Val	Ile	

Pro Ala Gly Ser Gly Lys Ser Thr Leu Leu Arg Glu Gly Phe Pro Ser

	Asp Pro	Ile	['] Val	Tyr	Thr	Pro	Glu	Ser	Ile	Arg	Gly	Val	Glu	Tyr	His
5		130		•			135					140			
	Leu Asp	Ile	Thr	Pro	Arg	Val	Gly	Asn	Gln	Ala	Val	Ile	Phe	Asp	Val
10	145 160	٠				150					155	,			
15	Gly Leu	Val	Leu	Thr	Thr	Pro	Gly	Gly	Asp	Asp	Leu	Leu	Arg	Arg	Arg
					165		+	•	•	170					175
20	Arg Pro	Glu	His	Trp	Leu	Gly	Trp	Leu	Met	Gln	Thr	Arg	Ala	Arg	Gln
				180					185					190	
25	Leu	Asn	Gly	Leu	Ile	Ļeu	Thr	Leu	Asp	Ľeu	Pro	.Asp	Leu	Leu	Thr
25	Ala		195					200					205		
	Asp	Lys	Ser	Arg	Arg	Glu	Thr	Leu	Val	Gln	Asn	Leu	Arg	Gln	Gln
30	Leu	210					215					220			
	Gln	Glu	Ile	Arg	Gln	Ser	Leu	His	Cys	Arg	Leu	Pro	Val	Tyr	Val
35	Val 225					230				•	235			-	
	240														,
40	Leu Ser	Thr	Arg	Leu	Asp	Leu	Leu	Asn	Gl _. y	Phe	Ala	Ala	Leu	Phe	His
,					245					250					255
45	Leu Arg	Asp	Lys	Lys	Asp	Arg	Asp	Ala	Ile	Leu	Gly	Val	Thr	Phe	Thr
	9			260		,			265			٠		270	
50	Arg Trp	Ala	His	Glu	Ser	Asp	Gly	Trp	Arg	Ser	Glu	Leu	Gly	Ala	Phe

			275					280					285		
. 5	Gln Leu	Thr 290	Trp	Val	Gln	Gln	Val 295	Asn	Leu	Ala	Leu	Ser	Asp	Leu	Val
10	Ala Arg 305 320	Gln	Thr	Gly	Ala ·	Ala	Pro	Arg	Ser	Ala	Val	Phe	Ser	Phe	Ser
15	Gln Leu	Met	Gln	Gly	Thr 325	Gly	Glu	Ile	Val	Thr 330	Ala	Leu	Leu	Ala	Ala
20	Leu Thr	Asp	Gly	Glu 340	Asn	Met	-Asp	Val	Met 345		Arg	Gly	Val	Trp 350	Leu
25	Ser Ala	Ser	Leu 355	Gln	Arg	Gly	Gln	Val 360	Asp	Asp	Ile	Phe	·Thr 365	Gln	Ser
30	Ala Leu	Arg 370	Gln	Tyr	Gly	Leu	Gly 375	Asn	Ser	Ser	Leu	Ala 380	Thr	Trp	Pro
35	Val Leu 385 400	Glu	Thr	Thr	Pro	Tyr 390	Phe	Thr	Arg	Arg	Leu 395	Phe	Pro	Glu	Val
40	Leu Ser	Ala	·Glu	Pro	Asn 405		Ala	Gly	Glu	Asn 410	Ser	Val	Trp	Leu	Asn 415
45	Ser Ala	Arg	Arg	Arg 420	Leu	Thr	Ala	Phe	Ser 425	Thr	Cys	Gly	Ala	Ala 430	Leu

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	Ala Gln	Leu	Met	.Val	Gly	Ser	Trp	His	His	Tyr	Tyr	Asn	Gln	Asn	Trp
	- 0111		435					440					445		
. 5		Gly	Val	Asn	Val	Leu	Ala	Gln	Ala	Lys	Ala	Phe	Met	Asp	Val
	Pro	450					455		,			4.60			
10	Pro	Pro	Gln	Glv	Thr	Asn	Glu	Phe	Glv	Asn	T. 2 11	Gln	Lou	Dro	T 011
	Leu 465	110	0111	· ·	1111	470	Giu	1110	О±у	ASII	475	GIII	ъеи	PLO	ьеи
15	480		٠								1.0				
	Asn His	Pro	Val	Àrg	Asp	Ala	Thr	Leu	Ala	Tyr	Gly	Asp	Tyr	Arg	Asp
20					485					490					495
		Phe	Leu	Ala	Asp	Met	Gly	Leu	Tyr	Gln	Gly	Ala	Arg	Val	Gly
25	Pro		•	500					505					510	
25	Trans.	77-1	Clu	Cln	mb s		т 1 -	<i>C</i> 1 m	Τ	T	G1	G1	70		
	Pro	val	Glu 515	GTII	TIIL	тйг	тте	520	ьеи	ьеи	GIU	GIN	Arg 525	Tyr	Leu
30	•		,					320					323		
	Ser Glu	Leu	Met	Asn	Gly	Leu	Ile	Arg	Asp	Leu	Asn	Ile	Ala	Pro	Pro
35		530		•		•	535					540			•
		Glu	Glu	Lys	Leu	Ala	Val	Leu	Arg	Val	Val	Arg	Met	Met	Glu
40	Asp 545 560					550			•		555				
10	300										•				
	Lys Arg	Ser	Gly	Arg	Asn	Asn	Glu	Ala	Val	Lys	Gln	Tyr	Met	Ala	Arg
45	2		-		565					570					575
		Ser	Asn	Glu	Phe	His	Gly	Gln	Arg	Asp	Ile	Gln	Ala	Gln	Leu
50	Met		•	580	•				585	,				590	

- 6							342/370								
5	Val Arg	His	Leu 595	Asp	Tyr	Ala	Leu	Glu 600	His	Thr	Asp	Trp	His	Ala	Gln

Gln Ser Ser Asp Ser Asp Ala Val Ser Arg Trp Thr Pro Tyr Asp Lys 610 615 620

10

15

Pro Ile Ile Asn Ala Gln Gln Glu Leu Ser Lys Leu Pro Ile Tyr Gln . 625 630 635

Arg Val Tyr Gln Thr Leu Arg Thr Lys Ala Leu Ser Val Leu Pro Ala . 645 650 655

Asp Leu Asn Leu Arg Asp Gln Val Gly Pro Thr Phe Asp Asn Val Phe
25 660 665 670

Val Ala Gly Asn Asp Glu Lys Leu Val Ile Pro Gln Phe Leu Thr Arg
30 675 680 685

Tyr Gly Leu Gln Ser Tyr Phe Val Lys Gln Arg Glu Gly Leu Val Glu
35 690 695 700

Leu Thr Ala Leu Asp Ser Trp Val Leu Asn Leu Thr Gln Ser Val
Ala
40 705 720 715

Tyr Ser Glu Ala Asp Arg Glu Glu Ile Gln Arg His Ile Thr Glu
45 Gln 725 730 735

Tyr Ile Ser Asp Tyr Thr Ala Thr Trp Arg Ala Gly Met Asp Asn 50 Leu 740 745 750

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	T.e.11	.· Glv	His	Glu	Phe	Ala	Pro	Glu	7) en	Sar	7\] >	T. 611	Clu	Clu	Cln
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- 25 Asn Asp Leu Thr Phe Gly Asp Asp Gly Arg Val Leu Ile Arg Glu Asp 980 985 . 990
- 30 Ile Arg Gln Gln Leu Asp Thr Ala Gln Lys Ile Arg Asp Ile Phe Phe 995 1000 1005
- 35 Ser Gln Gln Asn Gly Leu Gly Ala Gln Phe Ala Val Glu Thr Val 1010 1020
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- Gln Leu Val Asp Tyr Ser Gln Gly Arg Asn Tyr Thr Ala His Leu 1040 1045 1050
 - Val Trp Pro Asn Asn Met Arg Glu Gly Asn Glu Ser Lys Leu Thr 1055 1060 1065

Leu Ile Gly Thr Ser Gly Arg Ala Pro Arg Ser Ile Ala Phe Ser

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